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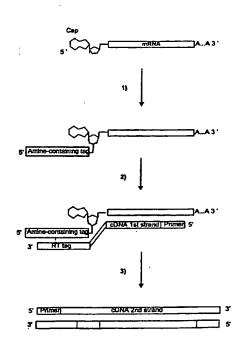
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(54) Title: 5' ESTS AND ENCODED HUMAN PROTEINS

(57) Abstract

The sequences of 5' ESTs derived from mRNAs encoding secreted proteins are disclosed. The 5' ESTs may be to obtain cDNAs and genomic DNAs corresponding to the 5' ESTs. The 5' ESTs may also be used in diagnostic, forensic, gene therapy, and chromosome mapping procedures. Upstream regulatory sequences may also be otained using the 5' ESTs. The 5' ESTs may also be used to design expression vectors and secretion vectors.



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	B. FIELDS SEARCHED									
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C DOCUM	ENTS CONSIDERED TO BE RELEVANT									
Category *	Citation of document, with indication, where appropriate, of the r	elevant passages	Relevant to claim No.							
X	BRENNER ET AL.: "Homo sapiens genomic DNA in the region of the locus" EMBL SEQUENCE DATABASE, 9 May 1996 (1996-05-09), XP0021	e L1CAM	1,2							
Υ	HEIDELBERG DE Ac U52112 the whole document & BRENNER ET AL.: "Genomic organization of two novel genes on human Xq28: compact head to head arrangement of IDH gamma and TRAP delta is conserved in rat and mouse" GENOMICS, vol. 44, no. 1, 1997, pages 8-14,									
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X Furth	er documents are listed in the continuation of box C.	X Patent family members	are listed in annex.							
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	COMMENTS CONFIDENCE TO DE SELEMANT	PC1/18 99/88/12
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Category ·	CALADI di CACAMITA NA STATEMENTO DE CACAMITA DE CACAMI	
X	SAKAI ET AL.: "Protein kinase C substrate, 80 kD protein, heavy chain (PKCSH)" SWISSPROT SEQUENCE DATA BASE, 1 January 1990 (1990-01-01), XP002121589	3
	HEIDELBERG DE Ac P14314 the whole document -& SAKAI ET AL.: "Human 80K-H protein (kinase C substrate) mRNA, complete compound" EMBL SEQUENCE DATABASE, 1 February 1990 (1990-02-01), XP002121590 HEIDELBERG DE Ac J03075 the whole document & SAKAI ET AL.: "Isolation of cDNAs encoding a substrate for protein kinase C: nucleotide sequence and chromosomal mapping of the gene for a human 80K protein" GENOMICS, vol. 5, 1989, pages 309-315,	
Y	WO 96 34981 A (GENSET ;MERENKOVA IRENA NICOLAEVNA (FR); DUMAS MILNE EDWARDS JEAN) 7 November 1996 (1996-11-07) cited in the application page 13, line 24 -page 14, line 14; claim 26	
Y	GREENWOOD M T ET AL: "Cloning of the gene encoding human somatostatin receptor 2: sequence analysis of the 50?-flanking promoter region" GENE, vol. 159, no. 2, 4 July 1995 (1995-07-04), page 291-292 XP004042228 ISSN: 0378-1119 abstract	5
Y	KATO S ET AL: "Construction of a human full-length cDNA bank" GENE, vol. 150, 1 January 1994 (1994-01-01), pages 243-250, XP602081364 ISSN: 0378-1119 cited in the application abstract page 245, left-hand column	6,10

S -- DOTAGE 2010 (comparison of second sheet) (July 19

INTERNATIONAL SEARCH REPORT

Intern. nat Application No PCT/IB 99/00712

	ation) DOCUMENTS CONSIDERED TO BE RELEVANT	- 1-7
Catogory *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Y	WO 97 38003 A (HUMAN GENOME SCIENCES INC; LI HAODONG (US); WEI YING FEI (US)) 16 October 1997 (1997-10-16) seg Id No 2	7,11,20, 21
n	claims 10-12	
Y	LOCKHART D J ET AL: "EXPRESSION MONITORING BY HYBRIDIZATION TO HIGH-DENSITY OLIGONUCLEOTIDE ARRAYS" BIO/TECHNOLOGY, vol. 14, no. 13, 1 December 1996 (1996-12-01), pages 1675-1680, XP002022521 ISSN: 0733-222X abstract	8,9 -
Υ	WO 98 07830 A (INST GENOMIC RESEARCH ;UNIV PENNSYLVANIA (US); UNIV JOHNS HOPKINS) 26 February 1998 (1998-02-26) page 3, line 4 - line 28 page 31, line 6 -page 35, line 16	7,11-21
X	MUZNY ET AL.: "Homo sapiens, working draft sequence, 97 unordered pieces" EMBL SEQUENCE DATABASE, 3 February 1998 (1998-02-03), XP002121591 HEIDELBERG DE AC AC004085 the whole document	1,2
x	ADAMS ET AL.: "EST177394 Jurkat T-cells VI homo sapiens cDNA 5' end similar to protein kinase C substrate 80K-H" EMBL SEQUENCE DATABASE, 18 April 1997 (1997-04-18), XP002121592 HEIDELBERG DE Ac AA306438 the whole document -& ADAMS ET AL.: "Initial assessment of human gene diversity and expression patterns based upon 83 million nucleotides of cDNA sequences" NATURE, vol. 377, 1995, pages 3-174, XP002069461	3
A	"zr94d07.rl NCI_CGAP_GCB1 Homo sapiens cDNA clone IMAGE:683341 5' EST" EMBL SEQUENCE DATABASE, 5 February 1997 (1997-02-05), XP002121593 HEIDELBERG DE Ac AA215334 the whole document	1,2
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INTERNATIONAL SEARCH REPORT

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C.(Continua	ation) DOCUMENTS CONSIDERED TO BE RELEVANT		
Category *	Citation of document, with indication, where appropriate, of the relevant passages	•	Relevant to claim No.
A	ADAMS M D ET AL: "RAPID CDNA SEQUENCING (EXPRESSED SEQUENCE TAGS) FROM A DIRECTIONALLY CLONED HUMAN INFANT BRAIN CDNA LIBRARY" NATURE GENETICS, vol. 4, no. 4, 1 August 1993 (1993-08-01), pages 373-380, STANDARD, XP002064427 ISSN: 1061-4036		_
А	ADAMS M D ET AL: "3,400 NEW EXPRESSED SEQUENCE TAGS IDENTIFY DIVERSITY OF TRANSCRIPTS IN HUMAN BRAIN" NATURE GENETICS, vol. 4, no. 3, 1 July 1993 (1993-07-01), pages 256-267, XP000645060 ISSN: 1061-4036		
A	TASHIRO K ET AL: "SIGNAL SEQUENCE TRAP: A CLONING STRATEGY FOR SECRETED PROTEINS AND TYPE I MEMBRANE PROTEINS" SCIENCE, vol. 261, 30 July 1993 (1993-07-30), pages 600-603, XP000673204 ISSN: 0036-8075		
A	CARNINCI P ET AL: "High-efficiency full-length cDNA cloning by biotinylated CAP trapper" GENOMICS, vol. 37, no. 3, 1 November 1996 (1996-11-01), pages 327-336, XP002081729 ISSN: 0888-7543		

International application No. PCT/IB 99/00712

INTERNATIONAL SEARCH REPORT

Box i	Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)
This int	ernational Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1. X	Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:
	Rule 39.1(v) PCT - Presentation of information Although claim 12 could be considered as a mere presentation of information, Rule 39.1(v) PCT, the search has been carried out as far as possible in our systematic documentation.
2.	Claims Nos.: because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically.
3.	Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II	Observations where unity of invention is lacking (Continuation of Item 2 of first sheet)
This Int	emational Searching Authority found multiple inventions in this international application, as follows:
1.	As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2.	As all searchable claims could be searched without effort justifying an additional tee, this Authority did not invite payment of any additional tee.
3.	As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
	i
4. X	No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:
	Invention 1: 1-21 partially
Remark	on Protest The additional search fees were accompanied by the applicant's protest.
	No protest accompanied the payment of additional search fees.

Form PCT/ISA/210 (continuation of first sheet (1)) (July 1998)

FURTHER INFORMATION CONTINUED FROM PCT/ISAV 210

Continuation of Box I.1

Although claim 12 could be considered as a mere presentation of information, Rule 39.1(v) PCT, the search has been carried out as far as possible in our systematic documentation.

Continuation of Box I.1

Rule 39.1(v) PCT - Presentation of information

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

1. Claims: Invention 1: 1-21 all partially

Nucleic acid comprising a sequence as in Seq.ID.No. 24, complementary sequence and fragments thereof. Polypeptide, Seq.Id.No. 812, encoded by said nucleotide sequence. Vector comprising Seq.Id.No. 24 and host cell comprising the vector. Methods of making cDNA and polypeptide utilising Seq.Id.No. 24. Array of ESTs comprising Seq.Id.No. 24, or a fragment thereof. An antibody binding to an epitop of the polypeptide of Seq.Id.No. 812. A computer readable medium and a computer system storing and/or utilising the sequence of Seq.Id.No. 24 or 812.

2. Claims: Invention 2-811 : 1-21 all partially

Idem as subject 1 but limited to each of the DNA sequences as in Seq.Id.No. 25-811 and 1600-1622, and corresponding polypeptides when applicible, where invention 2 is limited to Seq.Id.No. 25 and 813, invention 3 is limited to Seq.Id.No. 26 and 814,, invention 788 is limited to Seq.Id.No. 811 and 1599, invention 789 is limited to Seq.Id.No. 1600, invention 790 is limited to Seq.Id.No. 1601,, invention 811 is limited to Seq.Id.No. 1622.

INTERNATIONAL SEARCH REPORT

Information on patent family members

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Patent document cited in search report		Publication date		ratent family member(s)	Publication date
WO 9634981	A	07-11-1996	FR	2733765 A	08-11-1996
			FR	2733762 A	08-11-1996
			ΑU	5982996 A	21-11-1996
			CA	2220045 A	07-11-1996
			EP	0824598 A	25-02-1996
			JP	11510364 T	14-09-1999
WO 9738003	 A	16-10-1997	AU	5389096 A	29-10-1997
NO 3730003	••	20 20 200	ÜS	5945303 A	31-08-1999
WO 9807830	 A	26-02-1998	NONE		

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C12N 15/11, 15/10, C07K 14/47, C12P 21/00, C12Q 1/68, C07K 16/18, G06F 17/30, 17/50	A2	(4	3) International Publication Date: 21 October 1999 (21.10.99)
(21) International Application Number: PCT/IB (22) International Filing Date: 9 April 1999 ((81) Designated States: AU, CA, JP, US, European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE).
(30) Priority Data: 09/057,719 09/069,047 9 April 1998 (09.04.98) 28 April 1998 (28.04.98)		US US	Published Without international search report and to be republished upon receipt of that report.
(71) Applicant (for all designated States except US): [FR/FR]; 24, rue Royale, F-75008 Paris (FR).	GENSI	ET	·
(72) Inventors; and (75) Inventors/Applicants (for US only): DUMAS MIL WARDS, Jean-Baptiste [FR/FR]; 8, rue Grégoire- F-75006 Paris (FR). DUCLERT, Aymeric [FR/F] rue Victorine, F-94100 Saint-Maur (FR). GIO Jean-Yves [FR/FR]; 12, rue Duhesme, F-75018 Pa (74) Agents: MARTIN, Jean-Jacques et al.; Cabinet Re 26, avenue Kléber, F-75116 Paris (FR).	de–Tou R]; 6 t RDAN aris (FI	irs, ter, iO, R).	

(54) Title: 5' ESTS AND ENCODED HUMAN PROTEINS

(57) Abstract

The sequences of 5' ESTs derived from mRNAs encoding secreted proteins are disclosed. The 5' ESTs may be to obtain cDNAs and genomic DNAs corresponding to the 5' ESTs. The 5' ESTs may also be used in diagnostic, forensic, gene therapy, and chromosome mapping procedures. Upstream regulatory sequences may also be otained using the 5' ESTs. The 5' ESTs may also be used to design expression vectors and secretion vectors.

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5' ESTS AND ENCODED HUMAN PROTEINS

Background of the Invention

The estimated 50,000-100,000 genes scattered along the human chromosomes offer tremendous promise for the understanding, diagnosis, and treatment of human diseases. In addition, probes capable of specifically hybridizing to loci distributed throughout the human genome find applications in the construction of high resolution chromosome maps and in the identification of individuals.

In the past, the characterization of even a single human gene was a painstaking process, requiring years of effort. Recent developments in the areas of cloning vectors, DNA sequencing, and computer technology have merged to greatly accelerate the rate at which human genes can be isolated, sequenced, mapped, and characterized.

Currently, two different approaches are being pursued for identifying and characterizing the genes distributed along the human genome. In one approach, large fragments of genomic DNA are isolated, cloned, and sequenced. Potential open reading frames in these genomic sequences are identified using bioinformatics software. However, this approach entails sequencing large stretches of human DNA which do not encode proteins in order to find the protein encoding sequences scattered throughout the genome. In addition to requiring extensive sequencing, the bioinformatics software may mischaracterize the genomic sequences obtained, *i.e.*, labeling non-coding DNA as coding DNA and vice versa.

An alternative approach takes a more direct route to identifying and characterizing human genes. In this approach, complementary DNAs (cDNAs) are synthesized from isolated messenger RNAs (mRNAs) which encode human proteins. Using this approach, sequencing is only performed on DNA which is derived from protein coding portions of the genome. Often, only short stretches of the cDNAs are sequenced to obtain sequences called expressed sequence tags (ESTs). The ESTs may then be used to isolate or purify extended cDNAs which include sequences adjacent to the EST sequences. The extended cDNAs may contain all of the sequence of the EST which was used to obtain them or only a portion of the sequence of the EST which was used to obtain them. In addition, the extended cDNAs may contain the full coding sequence of the gene from which the EST was derived or, alternatively, the extended cDNAs may include portions of the coding sequence of the gene from which the EST was derived. It will be appreciated that there may be several extended cDNAs which include the EST sequence as a result of alternate splicing or the activity of alternative promoters. Alternatively, ESTs having partially overlapping sequences may be identified and configs comprising the consensus sequences of the overlapping ESTs may be identified.

In the past, these short EST sequences were often obtained from oligo-dT primed cDNA

libraries. Accordingly, they mainly corresponded to the 3' untranslated region of the mRNA. In part, the prevalence of EST sequences derived from the 3' end of the mRNA is a result of the fact that typical

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techniques for obtaining cDNAs, are not well suited for isolating cDNA sequences derived from the 5' ends of mRNAs (Adams et al., Nature 377:3-174, 1996, Hillier et al., Genome Res. 6:807-828, 1996).

In addition, in those reported instances where longer cDNA sequences have been obtained, the reported sequences typically correspond to coding sequences and do not include the full 5' untranslated 5 region (5'UTR) of the mRNA from which the cDNA is derived. Indeed, 5'UTRs have been shown to affect either the stability or translation of mRNAs. Thus, regulation of gene expression may be achieved through the use of alternative 5'UTRs as shown, for instance, for the translation of the tissue inhibitor of metalloprotease mRNA in mitogenically activated cells (Waterhouse et al, J Biol Chem. 265:5585-9. 1990). Furthermore, modification of 5'UTR through mutation, insertion or translocation events 10 may even be implied in pathogenesis. For instance, the fragile X syndrome, the most common cause of inherited mental retardation, is partly due to an insertion of multiple CGG trinucleotides in the 5'UTR of the fragile X mRNA resulting in the inhibition of protein synthesis via ribosome stalling (Feng et al, Science 268:731-4, 1995). An aberrant mutation in regions of the 5'UTR known to inhibit translation of the proto-oncogene c-myc was shown to result in upregulation of c-myc protein 15 levels in cells derived from patients with multiple myelomas (Willis et al, Curr Top Microbiol Immunol 224:269-76, 1997). In addition, the use of oligo-dT primed cDNA libraries does not allow the isolation of complete 5'UTRs since such incomplete sequences obtained by this process may not include the first exon of the mRNA, particularly in situations where the first exon is short. Furthermore, they may not include some exons, often short ones, which are located upstream of splicing sites. Thus, there 20 is a need to obtain sequences derived from the 5' ends of mRNAs.

While many sequences derived from human chromosomes have practical applications, approaches based on the identification and characterization of those chromosomal sequences which encode a protein product are particularly relevant to diagnostic and therapeutic uses. In some instances, the sequences used in such therapeutic or diagnostic techniques may be sequences which encode proteins which are secreted from the cell in which they are synthesized. Those sequences encoding secreted proteins as well as the secreted proteins themselves, are particularly valuable as potential therapeutic agents. Such proteins are often involved in cell to cell communication and may be responsible for producing a clinically relevant response in their target cells. In fact, several secretory proteins, including tissue plasminogen activator, G-CSF, GM-CSF, erythropoietin, human growth hormone, insulin, interferon-α, interferon-β, interferon-γ, and interleukin-2, are currently in clinical use. These proteins are used to treat a wide range of conditions, including acute myocardial infarction, acute ischemic stroke, anemia, diabetes, growth hormone deficiency, hepatitis, kidney carcinoma, chemotherapy-induced neutropenia and multiple sclerosis. For these reasons, extended cDNAs encoding secreted proteins or portions thereof represent a valuable source of therapeutic agents. Thus, there is a need for the

In addition to being therapeutically useful themselves, secretory proteins include short peptides, called signal peptides, at their amino termini which direct their secretion. These signal peptides are

encoded by the signal sequences located at the 5' ends of the coding sequences of genes encoding secreted proteins. These signal peptides can be used to direct the extracellular secretion of any protein to which they are operably linked. In addition, portions of the signal peptides called membranetranslocating sequences, may also be used to direct the intracellular import of a peptide or protein of 5 interest. This may prove beneficial in gene therapy strategies in which it is desired to deliver a particular gene product to cells other than the cells in which it is produced. Signal sequences encoding signal peptides also find application in simplifying protein purification techniques. In such applications, the extracellular secretion of the desired protein greatly facilitates purification by reducing the number of undesired proteins from which the desired protein must be selected. Thus, there exists a need to identify 10 and characterize the 5' portions of the genes for secretory proteins which encode signal peptides.

Sequences coding for non-secreted proteins may also find application as therapeutics or diagnostics. In particular, such sequences may be used to determine whether an individual is likely to express a detectable phenotype, such as a disease, as a consequence of a mutation in the coding sequence of a protein. In instances where the individual is at risk of suffering from a disease or other undesirable 15 phenotype as a result of a mutation in such a coding sequence, the undesirable phenotype may be corrected by introducing a normal coding sequence using gene therapy. Alternatively, if the undesirable phenotype results from overexpression of the protein encoded by the coding sequence, expression of the protein may be reduced using antisense or triple helix based strategies.

The secreted or non-secreted human polypeptides encoded by the coding sequences may also be 20 used as therapeutics by administering them directly to an individual having a condition, such as a disease, resulting from a mutation in the sequence encoding the polypeptide. In such an instance, the condition can be cured or ameliorated by administering the polypeptide to the individual.

In addition, the secreted or non-secreted human polypeptides or portions thereof may be used to generate antibodies useful in determining the tissue type or species of origin of a biological sample. The 25 antibodies may also be used to determine the cellular localization of the secreted or non-secreted human polypeptides or the cellular localization of polypeptides which have been fused to the human polypeptides. In addition, the antibodies may also be used in immunoaffinity chromatography techniques to isolate, purify, or enrich the human polypeptide or a target polypeptide which has been fused to the human polypeptide.

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Public information on the number of human genes for which the promoters and upstream regulatory regions have been identified and characterized is quite limited. In part, this may be due to the difficulty of isolating such regulatory sequences. Upstream regulatory sequences such as transcription factor binding sites are typically too short to be utilized as probes for isolating promoters from human genomic libraries. Recently, some approaches have been developed to isolate human promoters. One of 35 them consists of making a CpG island library (Cross et al., Nature Genetics 6: 236-244, 1994). The second consists of isolating human genomic DNA sequences containing SpeI binding sites by the use of SpeI binding protein. (Mortlock et al., Genome Res. 6:327-335, 1996). Both of these approaches have

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their limits due to a lack of specificity and of comprehensiveness. Thus, there exists a need to identify and systematically characterize the 5' portions of the genes.

The present 5' ESTs may be used to efficiently identify and isolate 5'UTRs and upstream regulatory regions which control the location, developmental stage, rate, and quantity of protein 5 synthesis, as well as the stability of the mRNA. Once identified and characterized, these regulatory regions may be utilized in gene therapy or protein purification schemes to obtain the desired amount and locations of protein synthesis or to inhibit, reduce, or prevent the synthesis of undesirable gene products.

In addition, ESTs containing the 5' ends of protein genes may include sequences useful as probes for chromosome mapping and the identification of individuals. Thus, there is a need to identify 10 and characterize the sequences upstream of the 5' coding sequences of genes.

Summary of the Invention

The present invention relates to purified, isolated, or enriched 5' ESTs which include sequences derived from the authentic 5' ends of their corresponding mRNAs. The term "corresponding mRNA" 15 refers to the mRNA which was the template for the cDNA synthesis which produced the 5' EST. These sequences will be referred to hereinafter as "5' ESTs." The present invention also includes purified, isolated or enriched nucleic acids comprising contigs assembled by determining a consensus sequences from a plurality of ESTs containing overlapping sequences. These contigs will be referred to herein as "consensus contigated 5'ESTs."

As used herein, the term "purified" does not require absolute purity; rather, it is intended as a relative definition. Individual 5' EST clones isolated from a cDNA library have been conventionally purified to electrophoretic homogeneity. The sequences obtained from these clones could not be obtained directly either from the library or from total human DNA. The cDNA clones are not naturally occurring as such, but rather are obtained via manipulation of a partially purified naturally occurring 25 substance (messenger RNA). The conversion of mRNA into a cDNA library involves the creation of a synthetic substance (cDNA) and pure individual cDNA clones can be isolated from the synthetic library by clonal selection. Thus, creating a cDNA library from messenger RNA and subsequently isolating individual clones from that library results in an approximately 104-106 fold purification of the native message. Purification of starting material or natural material to at least one order of magnitude, 30 preferably two or three orders, and more preferably four or five orders of magnitude is expressly contemplated.

As used herein, the term "isolated" requires that the material be removed from its original environment (e.g., the natural environment if it is naturally occurring). For example, a naturallyoccurring polynucleotide present in a living animal is not isolated; but the same polynucleotide, 35 separated from some or all of the coexisting materials in the natural system, is isolated.

As used herein, the term "recombinant" means that the 5' EST is adjacent to "backbone" nucleic acid to which it is not adjacent in its natural environment. Additionally, to be "enriched" the 5' ESTs will

represent 5% or more of the number of nucleic acid inserts in a population of nucleic acid backbone molecules. Backbone molecules according to the present invention include nucleic acids such as expression vectors, self-replicating nucleic acids, viruses, integrating nucleic acids, and other vectors or nucleic acids used to maintain or manipulate a nucleic acid insert of interest. Preferably, the enriched 5' ESTs represent 15% or more of the number of nucleic acid inserts in the population of recombinant backbone molecules. More preferably, the enriched 5' ESTs represent 50% or more of the number of nucleic acid inserts in the population of recombinant backbone molecules. In a highly preferred embodiment, the enriched 5' ESTs represent 90% or more of the number of nucleic acid inserts in the population of recombinant backbone molecules.

"Stringent," "moderate," and "low" hybridization conditions are as defined below.

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The term "polypeptide" refers to a polymer of amino acids without regard to the length of the polymer; thus, peptides, oligopeptides, and proteins are included within the definition of polypeptide. This term also does not specify or exclude post-expression modifications of polypeptides, for example, polypeptides which include the covalent attachment of glycosyl groups, acetyl groups, phosphate groups, lipid groups and the like are expressly encompassed by the term polypeptide. Also included within the definition are polypeptides which contain one or more analogs of an amino acid (including, for example, non-naturally occurring amino acids, amino acids which only occur naturally in an unrelated biological system, modified amino acids from mammalian systems etc.), polypeptides with substituted linkages, as well as other modifications known in the art, both naturally occurring and non-naturally occurring.

As used interchangeably herein, the terms "nucleic acids," "oligonucleotides," and "polynucleotides" include RNA, DNA, or RNA/DNA hybrid sequences of more than one nucleotide in either single chain or duplex form. The term "nucleotide" as used herein as an adjective to describe molecules comprising RNA, DNA, or RNA/DNA hybrid sequences of any length in single-stranded or duplex form. The term "nucleotide" is also used herein as a noun to refer to individual nucleotides or varieties of nucleotides, meaning a molecule, or individual unit in a larger nucleic acid molecule, comprising a purine or pyrimidine, a ribose or deoxyribose sugar moiety, and a phosphate group, or phosphodiester linkage in the case of nucleotides within an oligonucleotide or polynucleotide. Although the term "nucleotide" is also used herein to encompass "modified nucleotides" which comprise at least one modifications (a) an alternative linking group, (b) an analogous form of purine, (c) an analogous form of pyrimidine, or (d) an analogous sugar, for examples of analogous linking groups, purine, pyrimidines, and sugars see for example PCT publication No. WO 95/04064. The polynucleotide sequences of the invention may be prepared by any known method, including synthetic, recombinant, ex vivo generation, or a combination thereof, as well as utilizing any purification methods known in the art.

The terms "base paired" and "Watson & Crick base paired" are used interchangeably herein to refer to nucleotides which can be hydrogen bonded to one another be virtue of their sequence

identities in a manner like that found in double-helical DNA with thymine or uracil residues linked to adenine residues by two hydrogen bonds and cytosine and guanine residues linked by three hydrogen bonds (See Stryer, L., Biochemistry, 4th edition, 1995).

The terms "complementary" or "complement thereof" are used herein to refer to the 5 sequences of polynucleotides which are capable of forming Watson & Crick base pairing with another specified polynucleotide throughout the entirety of the complementary region. For the purpose of the present invention, a first polynucleotide is deemed to be complementary to a second polynucleotide when each base in the first polynucleotide is paired with its complementary base. Complementary bases are, generally, A and T (or A and U), or C and G. "Complement" is used 10 herein as a synonym from "complementary polynucleotide," "complementary nucleic acid" and "complementary nucleotide sequence". These terms are applied to pairs of polynucleotides based solely upon their sequences and not any particular set of conditions under which the two polynucleotides would actually bind. Preferably, a "complementary" sequence is a sequence which an A at each position where there is a T on the opposite strand, a T at each position where there is an A on 15 the opposite strand, a G at each position where there is a C on the opposite strand and a C at each position where there is a G on the opposite strand.

Thus, 5' ESTs in cDNA libraries in which one or more 5' ESTs make up 5% or more of the number of nucleic acid inserts in the backbone molecules are "enriched recombinant 5' ESTs" as defined herein. Likewise, 5' ESTs in a population of plasmids in which one or more 5' ESTs of the present 20 invention have been inserted such that they represent 5% or more of the number of inserts in the plasmid backbone are "enriched recombinant 5' ESTs" as defined herein. However, 5' ESTs in cDNA libraries in which 5' ESTs constitute less than 5% of the number of nucleic acid inserts in the population of backbone molecules, such as libraries in which backbone molecules having a 5' EST insert are extremely rare, are not "enriched recombinant 5' ESTs."

In some embodiments, the present invention relates to 5' ESTs which are derived from genes encoding secreted proteins. As used herein, a "secreted" protein is one which, when expressed in a suitable host cell, is transported across or through a membrane, including transport as a result of signal peptides in its amino acid sequence. "Secreted" proteins include without limitation proteins secreted wholly (e.g. soluble proteins), or partially (e.g. receptors) from the cell in which they are expressed. 30 "Secreted" proteins also include without limitation proteins which are transported across the membrane of the endoplasmic reticulum.

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Such 5' ESTs include nucleic acid sequences, called signal sequences, which encode signal peptides which direct the extracellular secretion of the proteins encoded by the genes from which the 5' ESTs are derived. Generally, the signal pentides are located at the amino termini of secreted proteins.

Secreted proteins are translated by ribosomes associated with the "rough" endoplasmic reticulum. Generally, secreted proteins are co-translationally transferred to the membrane of the endoplasmic reticulum. Association of the ribosome with the endoplasmic reticulum during translation WO 99/53051 PCT/IB99/00712

of secreted proteins is mediated by the signal peptide. The signal peptide is typically cleaved following its co-translational entry into the endoplasmic reticulum. After delivery to the endoplasmic reticulum, secreted proteins may proceed through the Golgi apparatus. In the Golgi apparatus, the proteins may undergo post-translational modification before entering secretory vesicles which transport them across 5 the cell membrane.

The 5' ESTs of the present invention have several important applications. For example, they may be used to obtain and express cDNA clones which include the full protein coding sequences of the corresponding gene products, including the authentic translation start sites derived from the 5' ends of the coding sequences of the mRNAs from which the 5' ESTs are derived. These cDNAs will be referred 10 to hereinafter as "full-length cDNAs." These cDNAs may also include DNA derived from mRNA sequences upstream of the translation start site. The full-length cDNA sequences may be used to express the proteins corresponding to the 5' ESTs. As discussed above, secreted proteins and non-secreted proteins may be therapeutically important. Thus, the proteins expressed from the cDNAs may be useful in treating and controlling a variety of human conditions. The 5' ESTs may also be used to obtain the 15 corresponding genomic DNA. The term "corresponding genomic DNA" refers to the genomic DNA which encodes the mRNA from which the 5' EST was derived.

Alternatively, the 5' ESTs may be used to obtain and express extended cDNAs encoding portions of the protein. In the case of secreted proteins, the portions may comprise the signal peptides of the secreted proteins or the mature proteins generated when the signal peptide is cleaved off.

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The present invention includes isolated, purified, or enriched "EST-related nucleic acids." The terms "isolated," "purified" or "enriched" have the meanings provided above. As used herein, the term "EST-related nucleic acids" means the nucleic acids of SEQ ID NOs. 24-811 and 1600-1622, extended cDNAs obtainable using the nucleic acids of SEQ ID NOs. 24-811 and 1600-1622, full-length cDNAs obtainable using the nucleic acids of SEQ ID NOs. 24-811 and 1600-1622 or genomic DNAs obtainable 25 using the nucleic acids of SEQ ID NOs. 24-811 and 1600-1622. The present invention also includes the sequences complementary to the EST-related nucleic acids.

The present invention also includes isolated, purified, or enriched "fragments of EST-related nucleic acids." The terms "isolated," "purified" and "enriched" have the meanings described above. As used herein the term "fragments of EST-related nucleic acids" means fragments comprising at least 10, 30 12, 15, 18, 20, 23, 25, 28, 30, 35, 40, 50, 75, 100, 200, 300, 500, or 1000 consecutive nucleotides of the EST-related nucleic acids to the extent that fragments of these lengths are consistent with the lengths of the particular EST-related nucleic acids being referenced. In particular, fragments of EST-related nucleic acids refer to "polynucleotides described in Table II," "polynucleotides described in Table III," and "polynuplectides described in Table IV." The present invention also includes the sequences 35 complementary to the fragments of the EST-related nucleic acids.

The present invention also includes isolated, purified, or enriched "positional segments of ESTrelated nucleic acids." As used herein, the term "positional segments of EST-related nucleic acids"

includes segments comprising nucleotides 1-25, 26-50, 51-75, 76-100, 101-125, 126-150, 151-175, 176-200, 201-225, 226-250, 251-300, 301-325, 326-350, 351-375, 376-400, 401-425, 426-450, 451-475, 476-500, 501-525, 526-550, 551-575, 576-600 and 601-the terminal nucleotide of the EST-related nucleic acids to the extent that such nucleotide positions are consistent with the lengths of the particular 5 EST-related nucleic acids being referenced. The term "positional segments of EST-related nucleic acids also includes segments comprising nucleotides 1-50, 51-100, 101-150, 151-200, 201-250, 251-300, 301-350, 351-400, 401-450, 450-500, 501-550, 551-600 or 601-the terminal nucleotide of the EST-related nucleic acids to the extent that such nucleotide positions are consistent with the lengths of the particular EST-related nucleic acids being referenced. The term "positional segments of EST-related nucleic 10 acids" also includes segments comprising nucleotides 1-100, 101-200, 201-300, 301-400, 501-500, 500-600, or 601-the terminal nucleotide of the EST-related nucleic acids to the extent that such nucleotide positions are consistent with the lengths of the particular EST-related nucleic acids being referenced. In addition, the term "positional segments of EST-related nucleic acids" includes segments comprising nucleotides 1-200, 201-400, 400-600, or 601-the terminal nucleotide of the EST-related nucleic acids to 15 the extent that such nucleotide positions are consistent with the lengths of the particular EST-related nucleic acids being referenced. The present invention also includes the sequences complementary to the positional segments of EST-related nucleic acids.

The present invention also includes isolated, purified, or enriched "fragments of positional segments of EST-related nucleic acids." As used herein, the term "fragments of positional segments of EST-related nucleic acids" refers to fragments comprising at least 10, 15, 18, 20, 23, 25, 28, 30, 35, 40, 50, 75, 100, 150, or 200 consecutive nucleotides of the positional segments of EST-related nucleic acids. The present invention also includes the sequences complementary to the fragments of positional segments of EST-related nucleic acids.

The present invention also includes isolated or purified "EST-related polypeptides." As used

herein, the term "EST-related polypeptides" means the polypeptides encoded by the EST-related nucleic acids, including the polypeptides of SEQ ID NOs. 812-1599.

The present invention also includes isolated or purified "fragments of EST-related polypeptides." As used herein, the term "fragments of EST-related polypeptides" means fragments comprising at least 5, 10, 15, 20, 25, 30, 35, 40, 50, 75, 100, or 150 consecutive amino acids of an EST-related polypeptide to the extent that fragments of these lengths are consistent with the lengths of the particular EST-related polypeptides being referenced. In particular, fragments of EST-related polypeptides refer to polypeptides encoded by "polynucleotides described in Table II," "polynucleotides described in Table III," and "polynucleotides described in Table IV."

The present invention also includes isolated or purified "positional segments of EST-related polypeptides." As used herein, the term "positional segments of EST-related polypeptides" includes polypeptides comprising amino acid residues 1-25, 26-50, 51-75, 76-100, 101-125, 126-150, 151-175, 176-200, or 201-the C-terminal amino acid of the EST-related polypeptides to the extent that such amino

acid residues are consistent with the lengths of the particular EST-related polypeptides being referenced. The term "positional segments of EST-related polypeptides also includes segments comprising amino acid residues 1-50, 51-100, 101-150, 151-200 or 201-the C-terminal amino acid of the EST-related polypeptides to the extent that such amino acid residues are consistent with the lengths of the particular 5 EST-related polypeptides being referenced. The term "positional segments of EST-related polypeptides" also includes segments comprising amino acids 1-100 or 101-200 of the EST-related polypeptides to the extent that such amino acid residues are consistent with the lengths of particular EST-related polypeptides being referenced. In addition, the term "positional segments of EST-related polypeptides" includes segments comprising amino acid residues 1-200 or 201-the C-terminal amino acid of the EST-10 related polypeptides to the extent that amino acid residues are consistent with the lengths of the particular EST-related polypeptides being referenced.

The present invention also includes isolated or purified "fragments of positional segments of EST-related polypeptides." As used herein, the term "fragments of positional segments of EST-related polypeptides" means fragments comprising at least 5, 10, 15, 20, 25, 30, 35, 40, 50, 75, 100, or 150 15 consecutive amino acids of positional segments of EST-related polypeptides to the extent that fragments of these lengths are consistent with the lengths of the particular EST-related polypeptides being referenced.

The present invention also includes antibodies which specifically recognize the EST-related polypeptides, fragments of EST-related polypeptides, positional segments of EST-related polypeptides, 20 or fragments of positional segments of EST-related polypeptides. In the case of secreted proteins, such as those of SEQ ID NOs. 1554-1580 antibodies which specifically recognize the mature protein generated when the signal peptide is cleaved may also be obtained as described below. Similarly, antibodies which specifically recognize the signal peptides of SEQ ID NOs. 812-1516 or 1554-1580 may also be obtained.

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In some embodiments and in the case of secreted proteins, the EST-related nucleic acids, fragments of EST-related nucleic acids, positional segments of EST-related nucleic acids, or fragments of positional segments of nucleic acids include a signal sequence. In other embodiments, the ESTrelated nucleic acids, fragments of EST-related nucleic acids, positional segments of EST-related nucleic acids, or fragments of positional segments of nucleic acids may include the full coding sequence for the 30 protein or, in the case of secreted proteins, the full coding sequence of the mature protein (i.e. the protein generated when the signal polypeptide is cleaved off). In addition, the EST-related nucleic acids, fragments of EST-related nucleic acids, positional segments of EST-related nucleic acids, or fragments of positional segments of nucleic acids may include regulatory regions upstream of the translation start site or downstream of the stop codon which control the amount, location, or developmental stage of gene 35 expression.

As discussed above, both secreted and non-secreted human proteins may be therapeutically important. Thus, the proteins expressed from the EST-related nucleic acids, fragments of EST-related

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nucleic acids, positional segments of EST-related nucleic acids, or fragments of positional segments of nucleic acids may be useful in treating or controlling a variety of human conditions.

The EST-related nucleic acids, fragments of EST-related nucleic acids, positional segments of EST-related nucleic acids, or fragments of positional segments of nucleic acids may be used in forensic procedures to identify individuals or in diagnostic procedures to identify individuals having genetic diseases resulting from abnormal gene expression. In addition, the EST-related nucleic acids, fragments of EST-related nucleic acids, positional segments of EST-related nucleic acids, or fragments of positional segments of nucleic acids are useful for constructing a high resolution map of the human chromosomes.

The present invention also relates to secretion vectors capable of directing the secretion of a protein of interest. Such vectors may be used in gene therapy strategies in which it is desired to produce a gene product in one cell which is to be delivered to another location in the body. Secretion vectors may also facilitate the purification of desired proteins.

The present invention also relates to expression vectors capable of directing the expression of an 15 inserted gene in a desired spatial or temporal manner or at a desired level. Such vectors may include sequences upstream of the EST-related nucleic acids, fragments of EST-related nucleic acids, positional segments of EST-related nucleic acids, or fragments of positional segments of nucleic acids, such as promoters or upstream regulatory sequences.

The present invention also comprises fusion vectors for making chimeric polypeptides 20 comprising a first polypeptide and a second polypeptide. Such vectors are useful for determining the cellular localization of the chimeric polypeptides or for isolating, purifying or enriching the chimeric polypeptides.

The EST-related nucleic acids, fragments of EST-related nucleic acids, positional segments of EST-related nucleic acids, or fragments of positional segments of nucleic acids may also be used for 25 gene therapy to control or treat genetic diseases. In the case of secreted proteins, signal peptides may be fused to heterologous proteins to direct their extracellular secretion.

Bacterial clones containing Bluescript plasmids having inserts containing the sequence of the non-aligned 5'ESTs, also referred to as singletons, and sequences of the 5'ESTs which were aligned to yield consensus contigated 5' ESTs are presently stored at 80°C in 4% (v/v) glycerol in the inventor's 30 laboratories under internal designations. The non-aligned 5'ESTs are those which comprise a single EST from a single tissue in the listing of Table V. The inserts may be recovered from the stored materials by growing the appropriate clones on a suitable medium. The Bluescript DNA can then be isolated using plasmid isolation procedures familiar to those skilled in the art such as alkaline lysis minipreps of large scale alkaline lysis plasmid isolation procedures. If desired the plasmid DNA may be-35 further enriched by centrifugation on a cesium chloride gradient, size exclusion chromatography, or anion exchange chromatography. The plasmid DNA obtained using these procedures may then be manipulated using standard cloning techniques familiar to those skilled in the art. Alternatively, a PCR

can be performed with primers designed at both ends of the inserted EST-related nucleic acids, fragments of EST-related nucleic acids, positional segments of EST-related nucleic acids, or fragments of positional segments of nucleic acids. The PCR product which corresponds to the EST-related nucleic acids, fragments of EST-related nucleic acids, positional segments of EST-related nucleic acids, or fragments of positional segments of nucleic acids can then be manipulated using standard cloning techniques familiar to those skilled in the art.

One embodiment of the present invention is a purified nucleic acid comprising a sequence selected from the group consisting of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622 and sequences complementary to the sequences of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622.

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Another embodiment of the present invention is a purified nucleic acid comprising at least 10, 12, 15, 18, 20, 23, 25, 28, 30, 35, 40, 50, 75, 100, 200, 300, 500, or 1000 consecutive nucleotides, to the extent that fragments of these lengths are consistent with the specific sequence, of a sequence selected from the group consisting of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622 and sequences complementary to the sequences of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622.

A further embodiment of the present invention is a purified nucleic acid comprising the coding sequence of a sequence selected from the group consisting of SEQ ID NOs. 24-811.

Yet another embodiment of the present invention is a purified nucleic acid comprising the full coding sequences of a sequence selected from the group consisting of SEQ ID NOs. 766-792 wherein the full coding sequence comprises the sequence encoding the signal peptide and the sequence encoding the mature protein.

Still another embodiment of the present invention is a purified nucleic acid comprising a contiguous span of a sequence selected from the group consisting of SEQ ID NOs. 766-792 which encodes the mature protein.

Another embodiment of the present invention is a purified nucleic acid comprising a contiguous span of a sequence selected from the group consisting of SEQ ID NOs. 24-728 and 766-792 which encodes the signal peptide.

Another embodiment of the present invention is a purified nucleic acid encoding a polypeptide comprising a sequence selected from the group consisting of the sequences of SEQ ID NOs. 812-1599.

Another embodiment of the present invention is a purified nucleic acid encoding a polypeptide comprising a sequence selected from the group consisting of the sequences of SEQ ID NOs. 1554-1580.

Another embodiment of the present invention is a purified nucleic acid encoding a

polypeptide comprising a mature protein included in a sequence selected from the group consisting of
the sequences of SEQ ID NOs. 1554-1580.

Another embodiment of the present invention is a purified nucleic acid encoding a polypeptide comprising a signal peptide included in a sequence selected from the group consisting of the sequences of SEQ ID NOs. 812-1516 and 1554-1580.

Another embodiment of the present invention is a purified nucleic acid at least 30, 35, 40, 50, 575, 100, 200, 300, 500 or 1000 nucleotides in length which hybridizes under stringent conditions to a sequence selected from the group consisting of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622 and sequences complementary to the sequences of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622.

Another embodiment of the present invention is a purified or isolated polypeptide comprising a sequence selected from the group consisting of the sequences of SEQ ID NOs. 812-1599.

Another embodiment of the present invention is a purified or isolated polypeptide comprising a sequence selected from the group consisting of SEQ ID NOs. 1554-1580.

Another embodiment of the present invention is a purified or isolated polypeptide comprising a mature protein of a polypeptide selected from the group consisting of SEQ ID NOs. 1554-1580.

Another embodiment of the present invention is a purified or isolated polypeptide comprising a signal peptide of a sequence selected from the group consisting of the polypeptides of SEQ ID NOs. 812-1516 and 1554-1580.

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Another embodiment of the present invention is a purified or isolated polypeptide comprising at least 12, 15, 18, 20, 23, 25, 28, 30, 35, 40, 50, 75, 100, 200, 300, 500, or 1000 consecutive amino acids, to the extent that fragments of these lengths are consistent with the specific sequence, of a sequence selected from the group consisting of the sequences of SEQ ID NOs. 812-1599.

Another embodiment of the present invention is a method of making a cDNA comprising the steps of contacting a collection of mRNA molecules from human cells with a primer comprising at least 12, 15, 18, 20, 23, 25, 28, 30, 35, 40, or 50 consecutive nucleotides of a sequence selected from the group consisting of the sequences complementary to SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622, hybridizing said primer to an mRNA in said collection that encodes said protein reverse transcribing said hybridized primer to make a first cDNA strand from said mRNA, making a second cDNA strand complementary to said first cDNA strand and isolating the resulting cDNA encoding said protein comprising said first cDNA strand and said second cDNA strand.

Another embodiment of the present invention is a purified cDNA obtainable by the method of the preceding paragraph.

In one aspect of this embodiment, the cDNA encodes at least a portion of a human polypeptide.

Another embodiment of the present invention is a method of making a cDNA comprising the steps of obtaining a cDNA comprising a sequence selected from the group consisting of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622, contacting said cDNA with a detectable probe comprising at least 12, 15, 18, 20, 23, 25, 28, 30, 35, 40, or 50 consecutive nucleotides of a sequence

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selected from the group consisting of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622 and the sequences complementary to SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622 under conditions which permit said probe to hybridize to said cDNA, identifying a cDNA which hybridizes to said detectable probe, and isolating said cDNA which hybridizes to said probe.

Another embodiment of the present invention is a purified cDNA obtainable by the method of the preceding paragraph.

In one aspect of this embodiment, the cDNA encodes at least a portion of a human polypeptide.

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Another embodiment of the present invention is a method of making a cDNA comprising the steps of contacting a collection of mRNA molecules from human cells with a first primer capable of hybridizing to the polyA tail of said mRNA, hybridizing said first primer to said polyA tail, reverse transcribing said mRNA to make a first cDNA strand, making a second cDNA strand complementary to said first cDNA strand using at least one primer comprising at least 12, 15, 18, 20, 23, 25, 28, 30, 35, 40, or 50 consecutive nucleotides of a sequence selected from the group consisting of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622, and isolating the resulting cDNA comprising said first cDNA strand and said second cDNA strand.

Another embodiment of the present invention is a purified cDNA obtainable by the method of the preceding paragraph.

In one aspect of this embodiment, said cDNA encodes at least a portion of a human 20 polypeptide.

In another aspect of the preceding method the second cDNA strand is made by contacting said first cDNA strand with a first pair of primers, said first pair of primers comprising a second primer comprising at least 12, 15, 18, 20, 23, 25, 28, 30, 35, 40, or 50 consecutive nucleotides of a sequence selected from the group consisting of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622 and a third primer having a sequence therein which is included within the sequence of said first primer, performing a first polymerase chain reaction with said first pair of primers to generate a first PCR product, contacting said first PCR product with a second pair of primers, said second pair of primers comprising a fourth primer, said fourth primer comprising at least 12, 15, 18, 20, 23, 25, 28, 30, 35, 40, or 50 consecutive nucleotides of said sequence selected from the group consisting of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622, and a fifth primer, wherein said fourth and fifth hybridize to sequences within said first PCR product, and performing a second polymerase chain reaction, thereby generating a second PCR product.

One aspect of this embodiment is a purified cDNA obtainable by the method of the preceding paragraph.

In another aspect of this embodiment, said cDNA encodes at least a portion of a human polypeptide.

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Alternatively, the second cDNA strand may be made by contacting said first cDNA strand with a second primer comprising at least 12, 15, 18, 20, 23, 25, 28, 30, 35, 40, or 50 consecutive nucleotides of a sequence selected from the group consisting of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622, hybridizing said second primer to said first strand cDNA, and extending said 5 hybridized second primer to generate said second cDNA strand.

One aspect of the above embodiment is a purified cDNA obtainable by the method of the preceding paragraph.

In a further aspect of this embodiment said cDNA encodes at least a portion of a human polypeptide.

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sites.

Another embodiment of the present invention is a method of making a polypeptide comprising the steps of obtaining a cDNA which encodes a polypeptide encoded by a nucleic acid comprising a sequence selected from the group consisting of SEQ ID NOs. 24-811 or a cDNA which encodes a polypeptide comprising at least 6, 8, 10, 12, 15, 18, 20, 23, 25, 28, 30, 35, 40, or 50 consecutive amino acids of a polypeptide encoded by a sequence selected from the group consisting 15 of SEQ ID NOs. 24-811, inserting said cDNA in an expression vector such that said cDNA is operably linked to a promoter, introducing said expression vector into a host cell whereby said host cell produces the protein encoded by said cDNA, and isolating said protein.

Another aspect of this embodiment is an isolated protein obtainable by the method of the preceding paragraph.

Another embodiment of the present invention is a method of obtaining a promoter DNA comprising the steps of obtaining genomic DNA located upstream of a nucleic acid comprising a sequence selected from the group consisting of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622 and the sequences complementary to the sequences of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622, screening said genomic DNA to identify a promoter capable of directing transcription 25 initiation, and isolating said DNA comprising said identified promoter.

In one aspect of this embodiment, said obtaining step comprises walking from genomic DNA comprising a sequence selected from the group consisting of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622 and the sequences complementary to SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622. In another aspect of this embodiment, said screening step comprises inserting genomic DNA located 30 upstream of a sequence selected from the group consisting of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622 and the sequences complementary to SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622 into a promoter reporter vector. For example, said screening step may comprise identifying motifs in genomic DNA located upstream of a sequence selected from the group consisting of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622 and the sequences complementary to SEQ ID NOs. 35 24-811 and SEQ ID NOs. 1600-1622 which are transcription factor binding sites or transcription start

Another embodiment of the present invention is a isolated promoter obtainable by the method of the paragraph above.

Another embodiment of the present invention is the inclusion of at least one sequence selected from the group consisting of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622, the sequences complementary to the sequences of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622 and fragments comprising at least 12, 15, 18, 20, 23, 25, 28, 30, 35, 40, 50, or 100 consecutive nucleotides of said sequence in an array of discrete ESTs or fragments thereof of at least 12, 15, 18, 20, 23, 25, 28, 30, 35, 40, 50, or 100 nucleotides in length. In some aspects of this embodiment, the array includes at least two sequences selected from the group consisting of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622, the sequences complementary to the sequences of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622, and fragments comprising at least 12, 15, 18, 20, 23, 25, 28, 30, 35, 40, 50, or 100 consecutive nucleotides of said sequences. In another aspect of this embodiment, the array includes at least one, three, five, ten, fifteen, or twenty sequences selected from the group consisting of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622, the sequences complementary to the sequences of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622 and fragments comprising at least 12, 15, 18, 20, 23, 25, 28, 30, 35, 40, 50, or 100 consecutive nucleotides of said sequences.

Another embodiment of the present invention is an enriched population of recombinant nucleic acids, said recombinant nucleic acids comprising an insert nucleic acid and a backbone nucleic acid, wherein at least 0.01%, 0.05%, 0.1%, 0.5%, 1%, 2%, 5%, 10%, or 20% of said insert nucleic acids in said population comprise a sequence selected from the group consisting of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622 and the sequences complementary to SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622.

Another embodiment of the present invention is a purified or isolated antibody capable of specifically binding to a polypeptide comprising a sequence selected from the group consisting of SEQ ID NOs. 812-1599.

Another embodiment of the present invention is a purified or isolated antibody capable of specifically binding to a polypeptide comprising at least 6, 8, 10, 12, 15, 18, 20, 23, 25, 28, 30, 35, 40, or 50 consecutive amino acids of a sequence selected from the group consisting of SEQ ID NOs. 812-1599.

Yet, another embodiment of the present invention is an antibody composition capable of selectively binding to an epitope-containing fragment of a polypeptide comprising a contiguous span of at least 8, 10, 12, 15, 18, 20, 23, 25, 28, 30, 35, 40, or 50 amino acids of any of SEQ ID NOs. 812-1599, wherein said antibody is polyclonal or monoclonal.

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Another embodiment of the present invention is a computer readable medium having stored thereon a sequence selected from the group consisting of a nucleic acid code of SEQ ID NOs. 24-811 and 1600-1622 and a polypeptide code of SEQ ID NOs. 812-1599.

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Another embodiment of the present invention is a computer system comprising a processor and a data storage device wherein said data storage device has stored thereon a sequence selected from the group consisting of a nucleic acid code of SEQID NOs. 24-811 and 1600-1622 and a polypeptide code of SEQ ID NOs. 812-1599. In one aspect of this embodiment the computer system further comprises a sequence comparer and a data storage device having reference sequences stored thereon. For example, the sequence comparer may comprise a computer program which indicates polymorphisms. In another aspect of this embodiment, the computer system further comprises an identifier which identifies features in said sequence.

Another embodiment of the present invention is a method for comparing a first sequence to a reference sequence wherein said first sequence is selected from the group consisting of a nucleic acid code of SEQID NOs. 24-811 and 1600-1622 and a polypeptide code of SEQ ID NOs. 812-1599 comprising the steps of reading said first sequence and said reference sequence through use of a computer program which compares sequences and determining differences between said first sequence and said reference sequence with said computer program. In some aspects of this embodiment, said step of determining differences between the first sequence and the reference sequence comprises identifying polymorphisms.

Another embodiment of the present invention is a method for identifying a feature in a sequence selected from the group consisting of a nucleic acid code of SEQID NOs. 24-811 and 1600-1622 and a polypeptide code of SEQ ID NOs. 812-1599 comprising the steps of reading said sequence through the use of a computer program which identifies features in sequences and identifying features in said sequence with said computer program.

Another embodiment of the present invention is a vector comprising a nucleic acid according to any one of the nucleic acids described above.

Another embodiment of the present invention is a host cell containing the above vector.

Another embodiment of the present invention is a method of making any of the nucleic acids described above comprising the steps of introducing said nucleic acid into a host cell such that said nucleic acid is present in multiple copies in each host cell and isolating said nucleic acid from said host cell.

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Another embodiment of the present invention is a method of making a nucleic acid of any of the nucleic acids described above comprising the step of sequentially linking together the nucleotides in said nucleic acids.

Another embodiment of the present invention is a method of making any of the polypeptides described above wherein said polypeptides is 150 amino acids in length or less comprising the step of sequentially linking together the amino acids in said polypeptide.

Another embodiment of the present invention is a method of making any of the polypeptides described above wherein said polypeptides is 120 amino acids in length or less comprising the step of sequentially linking together the amino acids in said polypeptides.

Brief Description of the Drawings

Figure 1 is a summary of a procedure for obtaining cDNAs which have been selected to include the 5' ends of the mRNAs from which they derived. In the first step (1), the cap of intact mRNAs is oxidized to be chemically ligated to an oligonucleotide tag. In the second step (2), a reverse transcription is performed using random primers to generate a first cDNA strand. In the third step (3), mRNAs are eliminated and the second strand synthesis is carried out using a primer contained in the oligonucleotide tag.

Figure 2 is an analysis of the 43 amino terminal amino acids of all human SwissProt proteins to determine the frequency of false positives and false negatives using the techniques for signal peptide identification described herein.

Figure 3 summarizes a general method used to clone and sequence extended cDNAs containing sequences adjacent to 5'ESTs.

Figure 4 provides a schematic description of the promoters isolated and the way they are assembled with the corresponding 5' tags.

Figure 5 describes the transcription factor binding sites present in each of the promoters of Figure 4.

Figure 6 is a block diagram of an exemplary computer system.

Figure 7 is a flow diagram illustrating one embodiment of a process 200 for comparing a new nucleotide or protein sequence with a database of sequences in order to determine the homology levels between the new sequence and the sequences in the database.

Figure 8 is a flow diagram illustrating one embodiment of a process 250 in a computer for determining whether two sequences are homologous.

Figure 9 is a flow diagram illustrating one embodiment of an identifier process 300 for detecting the presence of a feature in a sequence.

Figure 10 is a table with all of the parameters that can be used for each step of extended cDNA analysis.

Detailed Description of the Preferred Embodiment

30 I. Obtaining 5'ESTs from cDNA libraries including the 5'Ends of their Corresponding mRNAs

The 5' ESTs of the present invention were obtained from cDNA libraries including cDNAs which include the 5'end of their corresponding mRNAs. The general method used to obtain such cDNA libraries is described in Examples 1 to 5.

EXAMPLE:

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Preparation of mRNA

Total human RNAs or polyA⁺ RNAs derived from 29 different tissues were respectively purchased from LABIMO and CLONTECH and used to generate 44 cDNA libraries as described below.

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The purchased RNA had been isolated from cells or tissues using acid guanidium thiocyanate-phenol-chloroform extraction (Chomczyniski and Sacchi, *Analytical Biochemistry* 162:156-159, 1987). PolyA⁺ RNA was isolated from total RNA (LABIMO) by two passes of oligo dT chromatography, as described by Aviv and Leder, *Proc. Natl. Acad. Sci. USA* 69:1408-1412, 1972) in order to eliminate ribosomal 5 RNA.

The quality and the integrity of the polyA+ RNAs were checked. Northern blots hybridized with a globin probe were used to confirm that the mRNAs were not degraded. Contamination of the polyA+ mRNAs by ribosomal sequences was checked using Northern blots and a probe derived from the sequence of the 28S rRNA. Preparations of mRNAs with less than 5% of rRNAs were used in library construction. To avoid constructing libraries with RNAs contaminated by exogenous sequences (prokaryotic or fungal), the presence of bacterial 16S ribosomal sequences or of two highly expressed fungal mRNAs was examined using PCR.

EXAMPLE 2

Methods for Obtaining mRNAs having Intact 5' Ends

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Following preparation of the mRNAs from various tissues as described above, selection of mRNA with intact 5' ends and specific attachment of an oligonucleotide tag to the 5' end of such mRNA was performed using either a chemical or enzymatic approach. Both techniques takes advantage of the presence of the "cap" structure, which characterizes the 5'end of intact mRNAs and which comprises a guanosine generally methylated once, at the 7 position. The chemical approach is illustrated in Figure 1.

The chemical modification approach involves the optional elimination of the 2', 3'-cis diol of the 3' terminal ribose, the oxidation of the 2', 3', -cis diol of the ribose linked to the cap of the 5' ends of the mRNAs into a dialdehyde, and the coupling of the such obtained dialdehyde to a derivatized oligonucleotide tag. Further detail regarding the chemical approaches for obtaining mRNAs having intact 5' ends are disclosed in International Application No. WO96/34981, published November 7, 1996.

The enzymatic approach for ligating the oligonucleotide tag to the 5' ends of mRNAs with intact 5' ends involves the removal of the phosphate groups present on the 5' ends of uncapped incomplete mRNAs, the subsequent decapping of mRNAs with intact 5' ends and the ligation of the phosphate present at the 5' end of the decapped mRNA to an oligonucleotide tag. Further detail regarding the enzymatic approaches for obtaining mRNAs having intact 5' ends are disclosed in Dumas Milne Edwards J.B. (Doctoral Thesis of Paris VI University, Le clonage des ADNc complets: difficultes et perspectives nouvelles. Apports pour l'etude de la regulation de l'expression de la tryptophane hydroxylase de rat, 20 Dec. 1993), EP0 625572 and Kato et al., Gene 150:243-250 (1994).

In either the chemical or the enzymatic approach, the oligonucleotide tag has a restriction.

35 enzyme site (e.g. EcoRI sites) therein to facilitate later cloning procedures. Following attachment of the oligonucleotide tag to the mRNA, the integrity of the mRNA was then examined by performing a Northern blot using a probe complementary to the oligonucleotide tag.

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EXAMPLE 3

cDNA Synthesis Using mRNA Templates Having Intact 5' Ends

For the mRNAs joined to oligonucleotide tags, first strand cDNA synthesis was performed using 5 a reverse transcriptase with random nonamers as primers. In order to protect internal EcoRI sites in the cDNA from digestion at later steps in the procedure, methylated dCTP was used for first strand synthesis. After removal of mRNA by an alkaline hydrolysis, the first strand of cDNA was precipitated using isopropanol in order to eliminate residual primers.

The second strand of the cDNA was synthesized with a Klenow fragment using a primer 10 corresponding to the 5'end of the ligated oligonucleotide. Methylated dCTP was also used for second strand synthesis in order to protect internal EcoRI sites in the cDNA from digestion during the cloning process.

EXAMPLE 4

Cloning of cDNAs derived from mRNA with intact 5' ends into BlueScript 15

Following second strand synthesis, the ends of the cDNA were blunted with T4 DNA polymerase (Biolabs) and the cDNA was digested with EcoRI. Since methylated dCTP was used during cDNA synthesis, the EcoRI site present in the tag was the only hemi-methylated site, hence the only site susceptible to EcoRI digestion. The cDNA was then size fractionated using exclusion chromatography 20 (AcA, Biosepra) and fractions corresponding to cDNAs of more than 150 bp were pooled and ethanol precipitated. The cDNA was directionally cloned into the SmaI and EcoRI ends of the phagemid pBlueScript vector (Stratagene). The ligation mixture was electroporated into bacteria and propagated under appropriate antibiotic selection.

EXAMPLE 5 25

Selection of Clones Having the Oligonucleotide Tag Attached Thereto

Clones containing the oligonucleotide tag attached were then selected as follows. The plasmid DNAs containing 5' EST libraries made as described above were purified (Qiagen). A positive selection of the tagged clones was performed as follows. Briefly, in this selection procedure, the plasmid DNA 30 was converted to single stranded DNA using gene II endonuclease of the phage F1 in combination with an exonuclease (Chang et al., Gene 127:95-8, 1993) such as exonuclease III or T7 gene 6 exonuclease. The resulting single stranded DNA was then purified using paramagnetic beads as described by Fry et al., Biotechniques, 13: 124-131, 1992. In this procedure, the single stranded DNA was hybridized with a biotimylated oligonucleotide having a sequence corresponding to the 3' end of the oligonucleotide tag. 35 Clones including a sequence complementary to the biotinylated oligonucleotide were captured by incubation with streptavidin coated magnetic beads followed by magnetic selection. After capture of the positive clones, the plasmid DNA was released from the magnetic beads and converted into double

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stranded DNA using a DNA polymerase such as the Thermosequenase obtained from Amersham Pharmacia Biotech. The double stranded DNA was then electroporated into bacteria. The percentage of positive clones having the 5' tag oligonucleotide was estimated using dot blot analysis to typically be between 90 and 98%.

Following electroporation, the libraries were ordered in 384-microtiter plates (MTP). A copy of the MTP was stored for future needs. Then the libraries were transferred into 96 MTP and sequenced as described below.

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EXAMPLE 6

Sequencing of Inserts in Selected Clones

Plasmid inserts were first amplified by PCR on PE-9600 thermocyclers (Perkin-Elmer, Applied Biosystems Division, Foster City, CA), using standard SETA-A and SETA-B primers (Genset SA), AmpliTagGold (Perkin-Elmer), dNTPs (Boehringer), buffer and cycling conditions as recommended by the Perkin-Elmer Corporation.

PCR products were then sequenced using automatic ABI Prism 377 sequencers (Perkin Elmer). Sequencing reactions were performed using PE 9600 thermocyclers with standard dye-primer chemistry and ThermoSequenase (Amersham Pharmacia Biotech). The primers used were either T7 or 21M13 (available from Genset SA) as appropriate. The primers were labeled with the JOE, FAM, ROX and TAMRA dyes. The dNTPs and ddNTPs used in the sequencing reactions were purchased from 20 Boehringer. Sequencing buffer, reagent concentrations and cycling conditions were as recommended by Amersham.

Following the sequencing reaction, the samples were precipitated with ethanol, resuspended in formamide loading buffer, and loaded on a standard 4% acrylamide gel. Electrophoresis was performed for 2.5 hours at 3000V on an ABI 377 sequencer, and the sequence data were collected and analyzed 25 using the ABI Prism DNA Sequencing Analysis Software, version 2.1.2.

EXAMPLE 7

Obtaining 5' ESTs from Extended cDNA libraries Obtained from mRNA with Intact 5' Ends

Alternatively, 5'ESTs may be isolated from other cDNA or genomic DNA libraries. Such cDNA or genomic DNA libraries may be obtained from a commercial source or made using other techniques familiar to those skilled in the art. One example of such cDNA library construction, a fulllength cDNA library, is as follows.

PolyA+ RNAs are prepared and their quality sheeked as described in Example 1. Then, the 35 caps at the 5' ends of the polyA+ RNAs are specifically joined to an oligonucleotide tag as described in Example 2. The oligonucleotide tag may contain a restriction site such as Eco RI to facilitate further

subcloning procedures. Northern blotting is then performed to check the size of mRNAs having the oligonucleotide tag attached thereto and to ensure that the mRNAs are actually tagged.

First strand synthesis is subsequently carried out for mRNAs joined to the oligonucleotide tag as described in Example 3 above except that the random nonamers are replaced by an oligo-dT primer. For instance, this oligo-dT primer may contain an internal tag of 4 nucleotides which is different from one tissue to the other. Following second strand synthesis using a primer contained in the oligonucleotide tag attached to the 5' end of mRNA, the blunt ends of the obtained double stranded full-length DNAs are modified into cohesive ends to facilitate subcloning. For example, the extremities of full-length cDNAs may be modified to allow subcloning into the Eco RI and Hind III sites of a Bluescript vector using the Eco RI site of the oligonucleotide tag and the addition of a Hind III adaptor to the 3' end of full-length cDNAs.

The full-length cDNAs are then separated into several fractions according to their sizes using techniques familiar to those skilled in the art. For example, electrophoretic separation may be applied in order to yield 3 or 6 different fractions. Following gel extraction and purification, the cDNA fractions are subcloned into appropriate vectors, such as Bluescript vectors, transformed into competent bacteria and propagated under appropriate antibiotic conditions. Subsequently, plasmids containing tagged full-length cDNAs are positively selected as described in Example 5.

The 5' end of full-length cDNAs isolated from such cDNA libraries may then be sequenced as described in Example 6 to yield 5'ESTs.

II. Computer Analysis of the Isolated 5' ESTs: Construction of the SignalTag™ Database

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The sequence data from the cDNA libraries made as described above were transferred to a

database, where quality control and validation steps were performed. A base-caller, working using a
Unix system, automatically flagged suspect peaks, taking into account the shape of the peaks, the interpeak resolution, and the noise level. The base-caller also performed an automatic trimming. Any stretch
of 25 or fewer bases having more than 4 suspect peaks was considered unreliable and was discarded.
Sequences corresponding to cloning vector or ligation oligonucleotides were automatically removed
from the EST sequences. However, the resulting EST sequences may contain 1 to 5 bases belonging to
the above mentioned sequences at their 5' end. If needed, these can easily be removed on a case to case
basis.

Following sequencing as described above, the sequences of the 5' ESTs were entered in a database for storage and manipulation as described below. Before searching the ESTs in the database for sequences of interest, ESTs derived from mRNAs which were not of interest were identified. Briefly, such undesired sequences may be of three types. First, contaminants of either endogenous (ribosomal RNAs, transfert RNAs, mitochondrial RNAs) or exogenous (prokaryotic RNAs and fungal RNAs) origins were identified. Second, uninformative sequences, namely redundant sequences, small sequences and highly degenerate sequences were identified. Third, repeated sequences (Alu, L1, THE

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and MER repeats, SSTR sequences or satellite, micro-satellite, or telomeric repeats) were identified and masked in further processing.

In order to determine the accuracy of the sequencing procedure as well as the efficiency of the 5' selection described above, the analyses described in Examples 8 and 9 respectively were performed 5 on 5'ESTs obtained from the database following the elimination of endogenous and exogenous contaminants and following the masking of repeats.

EXAMPLE 8

Measurement of Sequencing Accuracy by Comparison to Known Sequences

To further determine the accuracy of the sequencing procedure described in Example 6, the sequences of 5' ESTs derived from known sequences were identified and compared to the original known sequences. First, a FASTA analysis with overhangs shorter than 5 bp on both ends was conducted on the 5' ESTs to identify those matching an entry in the public human mRNA database available at the time of filing the priority applications. The 5' ESTs which matched a known human 15 mRNA were then realigned with their cognate mRNA and dynamic programming was used to include substitutions, insertions, and deletions in the list of "errors" which would be recognized. Errors occurring in the last 10 bases of the 5' EST sequences were ignored to avoid the inclusion of spurious cloning sites in the analysis of sequencing accuracy. This analysis revealed that the sequences incorporated in the database had an accuracy of more than 99.5%.

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EXAMPLE 9

Determination of Efficiency of 5' EST Selection

To determine the efficiency at which the above selection procedures isolated 5' ESTs which included sequences close to the 5' end of the mRNAs from which they derived, the sequences of the 25 ends of the 5' ESTs derived from the elongation factor 1 subunit α and ferritin heavy chain genes were compared to the known cDNA sequences of these genes. Since the transcription start sites of both genes are well characterized, they may be used to determine the percentage of derived 5' ESTs which included the authentic transcription start sites. For both genes, more than 95% of the obtained 5' ESTs actually included sequences close to or upstream of the 5' end of the corresponding mRNAs.

To extend the analysis of the reliability of the procedures for isolating 5' ESTs from ESTs in the database, a similar analysis was conducted using a database composed of human mRNA sequences extracted from GenBank database release 97 for comparison. The 5' ends of more than 85% of 5' ESTs derived from mRNAs included in the GenBank database were located close to the 5' ends of the known sequence. As some of the mRNA sequences available in the GenBank database are deduced from 35 genomic sequences, a 5' end matching with these sequences will be counted as an internal match. Thus, the method used here underestimates the yield of ESTs including the authentic 5' ends of their corresponding mRNAs.

EXAMPLE 10

Calculation of Novelty Indices for 5'EST Libraries

In order to evaluate the novelty of 5'EST libraries, the following analysis was performed. For each sequenced 5'EST library, the sequences were clustered by the 5' end. Each sequence in the library was compared to the others and the longest sequence found in the cluster was used as representative of the group. A novelty rate (NR) was then defined as: NR= 100 X (Number of new unique sequences found in the library/Total number of sequences from the library). Typically, novelty rating ranged between 10% and 41% depending on the tissue from which the 5'EST library was obtained. For most of the libraries, the random sequencing of 5' EST libraries was pursued until the novelty rate reached 20%.

EXAMPLE 11

Generation of Consensus Contigated 5' ESTs

Since the cDNA libraries made above include multiple 5' ESTs derived from the same mRNA, overlapping 5'ESTs may be assembled into continuous sequences. The following method describes how to efficiently align multiple 5'ESTs in order to yield not only consensus contigated 5'EST sequences for mRNAs derived from different genes but also consensus contigated 5'EST sequences for different mRNAs, so called variants, transcribed from the same gene such as alternatively spliced mRNAs.

The whole set of sequences was first partitioned into small clusters containing sequences
which exhibited perfect matches with each other on a given length and which derived from a small number of different genes. Some 5'EST sequences, so called singletons, were not aligned using this approach because they were not homologous to any other sequence.

Thereafter, all variants of a given gene were identified in each cluster using a proprietary software. 5'EST sequences belonging to the same variant were then contigated and consensus contigated 5'EST sequences generated for each variant. All consensus contigated 5' EST sequences were subsequently compared to the whole set of individual 5'EST sequences used to obtained them.

If desired, the consensus contigated 5'EST sequences may be verified by identifying clones in nucleic acid samples derived from biological tissues, such as cDNA libraries, which hybridize to the probes based on the sequences of the consensus contigated 5'ESTs using any methods described herein and sequencing those clones.

Application of this alignment method to a selected set of 5'ESTs free from endogenous contaminants and uninformative sequences, and following the masking of repeats, yielded consensus contigated 5'ESTs sequences or variants of clustered genes encompassing many individual 5'ESTs.

Both non aligned 5'ESTs, i.e. singletons, and consensus contigated 5'ESTs were then compared to already known sequences and those sequences matching human mRNA sequences were eliminated from further analysis.

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EXAMPLE 12

Identification of Open Reading Frames in 5' ESTs

Subsequently, consensus contigated 5'ESTs and 5'ESTs were screened to identify those having an open reading frame (ORF).

Such open reading frames were simply defined as uninterrupted nucleic acid sequences longer than 45 nucleotides and beginning with an ATG codon.

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Alternatively, the nucleic acid sequence was first divided into several subsequences which coding propensity was evaluated separately using one or several different methods known to those skilled in the art such as the evaluation of N-mer frequency and its variants (Fickett and Tung, 10 Nucleic Acids Res; 20:6441-50 (1992)) or the Average Mutual Information method (Grosse et al, International Conference on Intelligent Systems for Molecular Biology, Montreal, Canada. June 28-July 1, 1998). Each of the scores obtained by the techniques described above were then normalized by their distribution extremities and then fused using a neural network into a unique score that represents the coding probability of a given subsequence. The coding probability scores obtained for 15 each subsequence, thus the probability score profiles obtained for each reading frame, was then linked to the initiation codons present on the sequence. For each open reading frame, defined as a nucleic acid sequence beginning with an ATG codon, an ORF score was determined. Preferably, this score is the sum of the probability scores computed for each subsequence corresponding to the considered ORF in the correct reading frame corrected by a function that negatively accounts for 20 locally high score values and positively accounts for sustained high score values. The most probable ORF with the highest score was selected.

In some embodiments, nucleic acid sequences encoding an "incomplete ORF", as referred therein, namely an open reading frame in which a start codon has been identified but no stop codon has been identified, were obtained.

In other embodiments, nucleic acid sequences encoding a "complete ORF", as used therein, namely an open reading frame in which a start codon and a stop codon have been identified, are obtained.

In a preferred embodiment, open reading frames encoding polypeptides of at least 50 amino acids were obtained.

To confirm that the chosen ORF actually encodes a polypeptide, the consensus contigated 5'EST or 5'EST may be used to obtain an extended cDNA using any of the techniques described therein, and especially those described in Examples 19 and 20. Then, such obtained extended cDNAs may be screened for the most probable open reading frame using any of the techniques described thereis. The amino acid sequence of the ORF encoded by the consensus contigated 5'EST or 5'EST may then be 35 compared to the amino acid sequence of the ORF encoded by the extended cDNA using any of the algorithms and parameters described therein in order to determine whether the ORF encoded by the extended cDNA is basically the same as the one encoded by the consensus contigated 5'EST or 5'EST.

Alternatively, to confirm that the chosen ORF actually encodes a polypeptide, the consensus contigated 5'EST or 5'EST may be used to obtain an extended cDNA using any of the techniques described therein, and especially those described in Examples 19 and 20. Such an extended cDNA may then be inserted into an appropriate expression vector and used to express the polypeptide encoded by 5 the extended cDNA as described therein. The expressed polypeptide may be isolated, purified, or enriched as described therein. Several methods known to those skilled in the art may then be used to determine whether the expressed polypeptide is the one actually encoded by the chosen ORF, therein referred to an the expected polypeptide. Such methods are based on the determination of predictable features of the expressed polypeptide, including but not limited to its amino acid sequence, its size or its 10 charge, and the comparison of these features to those predicted for the expected polypeptide. following paragraphs present examples of such methods.

One of these methods consists in the determination of at least a portion of the amino acid sequence of the expressed polypeptide using any technique known to those skilled in the art. For example, the amino-terminal residues may be determined using techniques either based on Sanger's 15 technique of acid hydrolysis of a polypeptide which N-terminal residue has been covalently labeled or using techniques based on Edman degradation of polypeptides which N-terminal residues are sequentially labeled and cleaved from the polypeptide of interest. The amino acid sequence of the expressed polypeptide may then be compared to the one predicted for the expected polypeptide using any algorithm and parameters described therein.

Alternatively, the size of the expressed polypeptides may be determined using techniques familiar to those skilled in the art such as Coomassie blue or silver staining and subsequently compared to the size predicted for the expected polypeptide. Generally, the band corresponding to the expressed polypeptide will have a mobility near that expected based on the number of amino acids in the open reading frame of the extended cDNA. However, the band may have a mobility different than that 25 expected as a result of modifications such as glycosylation, ubiquitination, or enzymatic cleavage.

Alternatively, specific antibodies or antipeptides may be generated against the expected polypeptide as described in Example 34 and used to perform immunoblotting or immunoprecipitation studies against the expressed polypeptide. The presence of a band in samples from cells containing the expression vector with the extended cDNA which is absent in samples from cells containing the 30 expression vector encoding an irrelevant polypeptide indicates that the expected polypeptide or portion thereof is being expressed. Generally, the band corresponding to the expressed polypeptide will have a mobility near that expected based on the number of amino acids in the open reading frame of the extended cDNA. However, the band may have a mobility different than that expected as a result of modifications such as glycosylation, ubiquitination, or enzymatic cleavage

The 5'ESTs or consensus contigated 5'ESTs found to encode an ORF were then searched to identify potential signal motifs using slight modifications of the procedures disclosed in Von Heijne, Nucleic Acids Res. 14:4683-4690, 1986. Those sequences encoding a 15 amino acid long stretch with a score of at least 3.5 in the Von Heijne signal peptide identification matrix were considered to possess a signal sequence. Those nucleic acid sequences which match a known human mRNA or EST sequence and have a 5' end located downstream of the known 5' end, preferably by more than 20 nucleotides, were excluded from further analysis. The remaining nucleic acids having signal sequences therein were included in a database called SignalTagTM.

10 EXAMPLE 14

Confirmation of Accuracy of Identification of Potential Signal Sequences in 5' ESTs

The accuracy of the above procedure for identifying signal sequences encoding signal peptides was evaluated by applying the method to the 43 amino acids located at the N terminus of all human SwissProt proteins. The computed Von Heijne score for each protein was compared with the known characterization of the protein as being a secreted protein or a non-secreted protein. In this manner, the number of non-secreted proteins having a score higher than 3.5 (false positives) and the number of secreted proteins having a score lower than 3.5 (false negatives) could be calculated.

Using the results of the above analysis, the probability that a peptide encoded by the 5' region of the mRNA is in fact a genuine signal peptide based on its Von Heijne's score was calculated based on either the assumption that 10% of human proteins are secreted or the assumption that 20% of human proteins are secreted. The results of this analysis are shown in Figure 2.

Using the above method of identification of secretory proteins, 5' ESTs of the following polypeptides known to be secreted were obtained: human glucagon, gamma interferon induced monokine precursor, secreted cyclophilin-like protein, human pleiotropin, and human biotinidase precursor. Thus, the above method successfully identified those 5' ESTs which encode a signal peptide.

To confirm that the signal peptide encoded by the 5' ESTs or consensus contigated 5' ESTs actually functions as a signal peptide, the signal sequences from the 5' ESTs or consensus contigated 5' ESTs may be cloned into a vector designed for the identification of signal peptides. Such vectors are designed to confer the ability to grow in selective medium only to host cells containing a vector with an operably linked signal sequence. For example, to confirm that a 5' EST or consensus contigated 5' EST encodes a genuine signal peptide, the signal sequence of the 5' EST or consensus contigated 5' EST may be inserted upstream and in frame with a non-secreted form of the yeast invertase gene in signal peptide selection vectors such as those described in U.S. Patent No. 5,336,637. Growth of host cells containing signal sequence selection vectors with the correctly inserted 5' EST or consensus contigated 5' EST signal sequence confirms that the 5' EST or consensus contigated 5' EST or consensus contigated 5' EST signal sequence confirms that the 5' EST or consensus contigated 5' ESTs encodes a genuine signal peptide.

Alternatively, the presence of a signal peptide may be confirmed by cloning the extended cDNAs obtained using the ESTs or consensus contigated 5' ESTs into expression vectors such as pXT1 as described below, or by constructing promoter-signal sequence-reporter gene vectors which encode fusion proteins between the signal peptide and an assayable reporter protein. After introduction of these 5 vectors into a suitable host cell, such as COS cells or NIH 3T3 cells, the growth medium may be harvested and analyzed for the presence of the secreted protein. The medium from these cells is compared to the medium from control cells containing vectors lacking the signal sequence or extended cDNA insert to identify vectors which encode a functional signal peptide or an authentic secreted protein.

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EXAMPLE 15

Analysis of the Sequences of the Invention

The set of the nucleic acid sequences of the invention (SEQ ID NOs.24-811 and 1600-1622) was obtained as described in Example 11. Subsequently, the most probable open reading frame was 15 determined and signal sequences were searched, as described in Examples 12 and 13, for all sequences of the invention.

The nucleotide sequences of the SEO ID NOs. 24-811 and 1600-1622 and the polypeptides sequences encoded by SEQ ID NOs. 24-811 (i.e. polypeptide sequences of SEQ ID NOs. 812-1599) are provided in the appended sequence listing which structure is as follows.

SEQ ID NOs. 24-728 are nucleic acids having an incomplete ORF which encodes a signal peptide. The locations of the incomplete ORFs and sequences encoding signal peptides are listed in the accompanying Sequence Listing. In addition, the von Heijne score of the signal peptide computed as described in Example 13 is listed as the "score" in the accompanying Sequence Listing. The sequence of the signal-peptide is listed as "seq" in the accompanying Sequence Listing. The "/" in the signal peptide 25 sequence indicates the location where proteolytic cleavage of the signal peptide occurs to generate a mature protein.

SEQ ID NOs. 729-765 are nucleic acids having an incomplete ORF in which no sequence encoding a signal peptide has been identified to date. However, it remains possible that subsequent analysis will identify a sequence encoding a signal peptide in these nucleic acids. The locations of the 30 incomplete ORFs are listed in the accompanying Sequence Listing.

SEQ ID NOs. 766-792 are nucleic acids having a complete ORF which encodes a signal peptide. The locations of the complete ORFs and of the signal peptides, the von Heijne score of the signal peptide, the sequence of the signal-peptide and the proteolytic cleavage site are indicated as described above.

SEQ ID NOs. 793-811 are nucleic acids having a complete ORF in which no sequence encoding a signal peptide has been identified to date. However, it remains possible that subsequent analysis will

identify a sequence encoding a signal peptide in these nucleic acids. The locations of the complete ORFs are listed in the accompanying Sequence Listing.

SEQ ID NOs. 812-1516 are "incomplete polypeptide sequences" which include a signal peptide. "Incomplete polypeptide sequences" are polypeptide sequences encoded by nucleic acids in which a start 5 codon has been identified but no stop codon has been identified. These polypeptides are encoded by the nucleic acids of SEQ ID NOs. 24-728. The location of the signal peptide, the von Heijne score of the signal peptide, the sequence of the signal-peptide and the proteolytic cleavage site are indicated as described above.

SEQ ID NOs. 1517-1553 are incomplete polypeptide sequences in which no signal peptide has 10 been identified to date. However, it remains possible that subsequent analysis will identify a signal peptide in these polypeptides. These polypeptides are encoded by the nucleic acids of SEQ ID NOs. 729-765.

SEQ ID NOs. 1554-1580 are "complete polypeptide sequences" which include a signal peptide. "Complete polypeptide sequences" are polypeptide sequences encoded by nucleic acids in which a start 15 codon and a stop codon have been identified. These polypeptides are encoded by the nucleic acids of SEQ ID NOs. 766-792. The location of the signal peptide, the von Heijne score of the signal peptide, the sequence of the signal-peptide and the proteolytic cleavage site are indicated as described above..

SEQ ID NOs. 1581-1599 are complete polypeptide sequences in which no signal peptide has been identified to date. However, it remains possible that subsequent analysis will identify a signal 20 peptide in these polypeptides. These polypeptides are encoded by the nucleic acids of SEQ ID NOs.793-811.

SEQ ID NOs. 1600-1622 are nucleic acid sequences in which no open reading frame has been conclusively identified to date. However, it remains possible subsequent analysis will identify an open reading frame in these nucleic acids.

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In the accompanying Sequence Listing, all instances of the symbol "n" in the nucleic acid sequences mean that the nucleotide can be adenine, guanine, cytosine or thymine. In some instances the polypeptide sequences in the Sequence Listing contain the symbol "Xaa." These "Xaa" symbols indicate either (1) a residue which cannot be identified because of nucleotide sequence ambiguity or (2) a stop codon in the determined sequence where applicants believe one should not exist (if the sequence 30 were determined more accurately). In some instances, several possible identities of the unknown amino acids may be suggested by the genetic code.

In the case of secreted proteins, it should be noted that, in accordance with the regulations governing Sequence Listings, in the appended Sequence Listing, the full protein (i.e. the protein containing the signal peptide and the mature protein) extends from an amino acid residue having a 35 negative number through a positively numbered C-terminal amino acid residue. Thus, the first amino acid of the mature protein resulting from cleavage of the signal peptide is designated as amino acid

number 1, and the first amino acid of the signal peptide is designated with the appropriate negative number.

If one of the nucleic acid sequences of SEQ ID NOs. 24-811 and 1600-1622 are suspected of containing one or more incorrect or ambiguous nucleotides, the ambiguities can readily be resolved by resequencing a fragment containing the nucleotides to be evaluated. If one or more incorrect or ambiguous nucleotides are detected, the corrected sequences should be included in the clusters from which the sequences were isolated, and used to compute other consensus contigated sequences on which other ORFs would be identified. Nucleic acid fragments for resolving sequencing errors or ambiguities may be obtained from deposited clones or can be isolated using the techniques described herein.

Resolution of any such ambiguities or errors may be facilitated by using primers which hybridize to sequences located close to the ambiguous or erroneous sequences. For example, the primers may hybridize to sequences within 50-75 bases of the ambiguity or error. Upon resolution of an error or ambiguity, the corresponding corrections can be made in the protein sequences encoded by the DNA containing the error or ambiguity. The amino acid sequence of the protein encoded by a particular clone can also be determined by expression of the clone in a suitable host cell, collecting the protein, and determining its sequence.

In addition, if one of the sequences of SEQ ID NOs. 812-1599 is suspected of containing a truncated ORF as the result of a frameshift in the sequence, such frameshifting errors may be corrected by combining the following two approaches. The first one involves thorough examination of all double predictions, *i.e.* all cases where the probability scores for two ORFs located on different reading frames are high and close, preferably different by less than 0.4. The fine examination of the region where the two possible ORFs overlap may help to detect the frameshift. In the second approach, homologies with known proteins are used to correct suspected frameshifts.

Of the identified clusters, some were shown to be multivariant, *i.e.* to contain several variants of the same gene. Table I gives for each of the multivariant clusters named by its internal reference (first column), the list of all variant consensus contigated 5'ESTs (second column), each being represented by a different sequence identification number.

TABLE I

Cluster Internal Reference	SEQ ID NOs of Variants
Cl	687, 791
C2	744, 798
C3	640, 811
C4	59, 66
C5	84, 97

30	
C6	287, 289
C7	286, 775, 777
C8	762, 768
C9	783, 784
C10	80, 1603
C11	655, 736
C12	805, 806

Table II provides a list preferred polynucleotide fragments which are derivatives of the consensus contigated 5'ESTs. As used herein the term "polynucleotide described in Table II" refers to the all of the preferred polynucleotide fragments defined in Table II in the following manner. The fragments are referred to by their SEQ ID numbers in the first column. The preferred polynucleotide fragments are then defined by a range of nucleotide positions from the SEQ IDs of the consensus contigated 5'ESTs as indicated in the second column entitled "positions of preferred fragments." The preferred polynucleotide fragments correspond to the individual 5'ESTs aligned to obtain the consensus contigated 5'EST and to those filed in the priority documents. The third column entitled "variant nucleotides" describes the nucleotide sequence variations observed between the consensus contigated 5'EST and preferred nucleic acid fragments as follows:

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A) Substitutions in the sequence of a consensus contigated 5'EST to derive a preferred polynucleotide fragment are denoted by an "S", followed by a number indicating the first nucleotide position in a specific SEQ ID to be substituted in a string of substituted nucleotides or the position of the substituted nucleotide in the case of a single substituted nucleotide. Then there is a coma followed by one or more lower case letters indicating the identity of the nucleotide(s) occurring in the substituted position(s). For example, SEQ ID NO: 3401; Position of preferred fragments: 1-250; Variant nucleotides S45,atc would indicate that a preferred polynucleotide fragment had the sequence of positions 1 to 250 of SEQ ID NO. 3401, except that the nucleotides at positions 45, 46, and 47 were substituted with A, T, and C, respectively, in the preferred polynucleotide as compared with the sequence of SEQ ID No. 3401.

B) Insertions in the sequence of a consensus contigated 5'EST to derive a preferred polynucleotide fragment are denoted by an "I", followed by a number indicating the nucleotide position in a specific SEQ ID after which a string of nucleotides is inserted or the position after which the nucleotide is inserted in the case of a single inserted nucleotide. Then there is a coma followed by one or more lower case letters indicating the identity of the nucleotide(s) occurring in the inserted position(s). For example, SEQ ID NO: 7934; Position of preferred fragments: 1-500; Variant nucleotides: I36,gataca would indicate that a preferred polynucleotide fragment had the sequence of positions 1 to 500 of SEQ ID NO: 7934, except that after the nucleotides at position 36 a GATACA string of nucleotides is inserted in the preferred polynucleotide as compared with the sequence of SEQ ID No. 7934.

C) Deletions in the sequence of a consensus contigated 5'EST to derive a preferred nucleic acid fragment are denoted by an "D", followed by a number indicating the first nucleotide position in a specific SEQ ID to be deleted in a string of deleted nucleotides or the position of the deleted nucleotide in the case of a single deleted nucleotide. Then there is a coma followed by number indicating the number of nucleotide(s) deleted from the sequence provided in the sequence ID. For example, SEQ ID NO: 5398; Position of preferred fragments: 56-780; Variant nucleotides D114,5 would indicate that a preferred polynucleotide fragment had the sequence of positions 56 to 780 of SEQ ID NO. 5398, except that the nucleotides in positions 114 to 118 had been deleted in the preferred polynucleotide as compared with the sequence of SEQ ID No. 5398.

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The present invention encompasses isolated, purified, or recombinant nucleic acids which consist of, consist essentially of, or comprise a contiguous span of at least 8, 10, 12, 15, 18, 20, 25, 35, 40, 50, 70, 80, 100, 250, or 500 nucleotides in length, to the extent that a contiguous span of these lengths is consistent with the lengths of the particular polynucleotide, of a polynucleotide described in Table II, or a sequence complementary thereto, wherein said polynucleotide described in Table II is selected individually or in any combination from the polynucleotides described in Table II. The present invention also encompasses isolated, purified, or recombinant nucleic acids which consist of or consist essentially of a polynucleotide described in Table II, or a sequence complementary thereto, wherein said polynucleotide is selected individually or in any combination from the polynucleotides described in Table II. The present invention further encompasses isolated or purified polypeptides which consist of, consist essentially of, or comprise a contiguous span of at least 8, 10, 12, 15, 18, 20, 25, 35, 40, 50, 70, 80, or 100 amino acids encoded by a polynucleotide described in Table II.

Table II

SEQ ID NO.	Positions of Preferred Fragments	Variant nucleotides
35	1-423	S124, s; I135, a; S293, w; I363, a; S377, r; D424, 15
41	1-427	I117, m; S120, r; S124, g; D373, 1; S376, b; S378, b; I427, gggg; D428, 109
43	1-276	S114, m; S118, rg; S123, r; S139, nr; I142, t; D148, 1; D152, 1; I228, t; I276, gg; D277, 136
45	126-420	D1, 125; I420, ggg; D421, 100
46	1-255	S139, r; I145, r; S146, mm; S150, ar; S254, g; D256, 128
48	4-437	D1, 3; S49, a; S55, g; S79, a; S90, a; I437, teteto
59	1-471	S26, a; S44, t; S48, t; S109, a; S191, t; S200, gc; S203, a; S210, g; S237, a; S240, g; S255, a; S272, a; S277, a; S279, a; S284, t; S297, g; S305, g; S316, a; I471, ggtca
66	1-428	I428, tactgggg

		32
82	1-399	S251, t; S277, d; I399, aagccggg
84	5 -4 88	D1, 4; S210, g; S293, a; S325, g; S339, a;
		S348, g; S353, g; S395, g; I488, cacca
93	1-508	I508, gattt
96	26-315	D1, 25; S28, a; S62, c; I315, cagatgg
97	4-460	D1, 3; S19, g; S31, g; S114, gt; S118, a; S123,
		tc; S127, c; S132, a; S186, g; S190, c; S203, t;
		S210, g; S232, c; I460, acgtt
105	1-281	S273, a; I281, g; D282, 211
114	10-315	I0, t; D1, 9; S91, m; S267, n; S276, w; S292, h;
		S295, m; I315, tggg; D316, 19
118	1-145	S57, d; S126, d; I145, ccctc
120	2-348	D1, 1; S104, t; I348, g; D349, 38
121	1-190	I121, c; I190, ccctt
123	1-353	I117, m; I186, w; S187, y; I353, caccgggg
124	1-249	I249, ggrvgggg
125	114-375	D1, 113; S206, wn; I231, a; I375, ccctagg
126	1-437	S297, cc; S307, tg; S312, a; S318, g; S341, a;
		S351, t; S353, g; S383, c; S387, a; D404, 1
136	82-428	D1, 81; I428, aaagtg
139	1-268	I268, gggaaggg
148	6-405	D1, 5; I405, ggtgt
159	1-230	S227, ta; I230, ccctggg
165	3-256	I0, tat; D1, 2; I17, c; S18, t; S111, d; I115, t;
		S123, r; I256, aaggeggg
170	1-280	I103, t; S104, c; I111, t; I280, cgttcggg
194	1-215	S50, s; S186, sn; S199, k; I215, gcagcggg
213	1-158	S128, m; I132, w; S143, d; I158, tgcccggg
223	3-431	D1, 2; S28, s; S79, c; S82, s; S308, nr; S328,
		nb; I431, ccggc
247	1-359	I76, gttt; I359, tccctgg
258	1-236	S72, r; S81, g; S197, s; I205, ss; S232, k; I236, actteggg
264	5-283	D1, 4; S64, g; S122, m; S134, yy; I137, c;
207	3-203	I151, t; I283, gttgc_
269	1-143	S111, s; I143, ggggcggg
286	5-207	D1, 4; S204, a; S206, c; I207, gg; D208, 567
287	1-277	S114, r; I125, t; S131, ag; S256, tg; S259, tt;
] =="		S262, at; S267, t; S269, c; S273, c; I277,
		ccggg; D278, 337
289	69-416	D1, 68; I416, agccaggg
289	1-278	S114, r; I125, t; S131, ag; S277, c; I278, cggg;
	ļ	D279, 138
292	20-254	D1, 19; I254, aaagagg
293	1-414	I414, tagcag
300	1-285	S16, m; S67, y; I285, baccacggg; D286, 1
349	23-431	Dî, 22; Iì 18, a; S214, y; I431, caactgg
350	3-386	D1, 2; S42, w; I263, c; I386, gggat
368	3-446	D1, 2; I446, tetet
385	1-193	I35, t; I108, t; I134, r; S135, a; S137, r; S143,
1		w; I178, c; I193, gagcgggg
411	6-391	D1, 5; S17, r; S27, t; S334, y; D392, 244
412	1-185	S49, s; S127, s; I185, gctggg; D186, 150
		. — — · — · — · — · — · — · — · — · — ·

		33
415	2-229	D1, 1; S3, a; I229, caaatggg
435	1-386	S4, s; I386, ccggg
436	4-472	D1, 3; S61, sa; D238, 1; S239, s; I472, agtgtgg
437	1-340	I340, ggg; D341, 129
441	1-409	S109, smag; I409, cgcacggg
454	1-492	S72, nn; S115, t; S121, bwy; S181, yn; I492,
		gagtc
455	1-177	I14, w; I16, a; I177, gagctggg
459	1-311	S39, n; S74, rg; I311, accatggg
460	1-425	I425, agtac
461	5-420	D1, 4; I420, tcgtc
481	1-429	I10, w; S262, d; S333, n; I429, ctccaggg
489	1-414	D72, 1; S117, n; S396, d; I414, ggaca
496	1-215	1215, ttttcggg
501	1-430	S275, n; I430, aggat
	91-413	D1, 90; I413, aaacgggg
502		D1, 20; S47, w; S83, n; I280, n; S281, na;
504	21-420	S292, v; S314, sm; S368, ww; S373, w; I420,
		cccca
505	18-457	D1, 17; D36, 1; S182, g; S273, n; S283, a;
505	10-43/	S416, bh; I457, ctcga
514	1-303	
514	1-303	I303, accca S11, t; I12, n; S30, r; S256, wr; I333, t; I455,
515	1-455	cataa
517	24-453	
517		D1, 23; I453, agagcggg I119, gt; S125, w; I129, w; S133, k; S137, k;
519	1-275	S167, k; I275, gcccc
	1-313	
522	4-366	I313, agcgtggg
526		I0, t; D1, 3; I366, ggcccggg
530	1-434	S328, g; I434, aagat S128, g; S162, m; D380, 5
535	1-379	
561	2-341	D1, 1; I341, raagagg
568	1-246	I118, g; S137, g; I246, aaaccggg
570	1-207	1207, ttttt
576	1-288	134, c; 1288, cccgtgg
588	1-390	S218, a; S224, k; S314, dh; S358, s; D376, 1;
	21.074	1390, atg; D391, 23
597	31-274	D1, 30; S49, n; I274, tccatgg
606	1-354	[1141, g; D174, 1; S229, п; D355, 72
627	1-415	S7, a; I415, cattt
634	1-178	D179, 212
640	6-428	D1, 5; D429, 79
641	64-483	D1, 63; 1165, d; D183, 1; S185, y; S253, t;
		D279, 2; S416, a; I483, atata
655	1-280	S58, c; I84, g; S88, k; S204, ac; S244, g; S247,
	<u> </u>	g; I280, ggg; D281, 90
672	34-489	D1, 33; S316, k; S331, k; S333, w; S486, g;
<u> </u>	<u> </u>	- S488, c; D490, 4
687	116-473	D1, 115; S142, n; I473, cctcgggg
697	1-202	S142, s; S144, sr; S148, d; S152, d; I155, a;
		I164, a; S174, k; I202, gcc; D203, 291
708	8-384	D1, 7; S104, b; I384, gaaaa
710	1-167	S40, k; S49, db; I167, tatct

722	1-191	I125, c; I191, tttt
723	1-316	I316, aggg; D317, 157
729	15-373	D1, 14; S139, t; I373, cgcag; D374, 99
730	29-372	D1, 28; I155, g; S192, ka; S333, d; I372, m;
730	25-372	D373, 93
731	1-290	S10, kk; S30, b; S32, t; S92, t; S197, dy; S278,
		g; I290, aggg; D291, 55
732	8-277	D1, 7; I113, a; S127, w; I131, s; S132, r; S156,
		w; S160, r; S211, n; S215, w; I247, a; D278,
		121
733	20-375	D1, 19; S306, sbs; I325, h; S326, nr; S338,
		ywd; S344, v; I375, aggg; D376, 68
734	1-359	D66, 1; D360, 14
735	25-322	D1, 24; S30, r; I193, a; I322, ccaaggg
736	9-181	D1, 8; S58, g; I181, aactaggg
737	1-160	S97, ta; I160, aggtc
738	1-227	D228, 7
739	45-514	D1, 44; S178, s; I182, c; S436, dmn; S461, v;
		S476, c; S506, t; D515, 75
740	11-388	D1, 10; I388, cgacaggg
741	1-478	S118, s; S125, a; I126, s; S134, k; S421, vn;
		1478, aatsc
742	217-553	10, tt; D1, 216; S286, r; S294, m; S311, r;
		S317, s; S338, r; S442, dm; S469, h; S476, r;
·		S485, s; S491, w; I495, ht; S496, v; S513, r;
742	1 450	D521, 1; S536, m; D554, 199
743	1-459	II1, s; S258, m; I270, m; I304, c; I308, amta;
744	25-316	S313, c; S438, v; I459, agggag D1, 24; S315, g; D317, 95
745	21-283	D1, 20; I40, g; S41, c; D123, 1; S181, sr; S227,
143	21-203	r; I283, ccgcg; D284, 121
746	1-256	D257, 173
747	1-179	S134, w; S138, w; S140, kt; I179, cacca
748	1-235	S46, t; I72, t; S189, cc; S222, c; D236, 148
749	2-370	D1, 1; S32, cg; D144, 1; S341, g; D371, 76
750	18-410	I0, aag; D1, 17; I410, aatcc
751	22-355	D1, 21; D148, 1; S150, c; S152, a; S313, n;
		D356, 181
752	1-139	S50, t; I118, g; I139, ccct
753	1-189	S26, r; S115, s; I121, r; S122, r; S128, s; S143,
		r; I146, w; S156, r; D190, 4
754	1-395	S212, wd; I395, cggca
755	19-460	D1, 18; S26, c; S156, a; S253, n; I460, tagaagg
756	2-142	D1, 1; I106, gc; S107, t; S110, c; I142,
		ccaccggg
757	28-296	D1, 27; I119, s; I122, t; S128, s; S255, t; S267,
<u> </u>	1	m; D297, 66
758	11-368	D1, 10; I200, g; S201, c; S281, d; S317, c;
	1	I368, ccateggg
759	19-452	D1, 18; S421, w; I452, a
760	25-175	D1, 24; S34, yk; I175, ccggg; D176, 120
761	1-212	I212, cacteggg
762 763	1-374	S320, s; S349, a; D375, 249
	8-152	D1, 7; I152, acggg; D153, 109

764			35
D314, 203 S278, ag; S281, cagacc; S288, ta; S291, cagg; S296, c; S317, m; I320, cggg; D321, 306 767	764	1-160	I127, g; I145, g; I160, cgcccggg
Total	765	137-313	
S296, c; S317, m; I320, eggg; D321, 306			
767 6-336 10, aa; D1, 5; S149, w; S245, y; D337, 137 768 1-374 320, s; D375, 299 769 53-435 D1, 52; S59, b; S344, nnkw; D436, 104 770 24-448 D1, 23; S25, g; S411, w; S416, m; D449, 31 771 1-370 S3, c; S180, m; S275, r; D371, 122 772 1-388 1299, c; S326, c; D389, 8 773 1-143 S18, c; S66, a; I143, ggg; D144, 274 774 1-347 S194, a; S205, c; I347, ggg; D348, 107 775 5-207 D1, 4; S111, tg; S158, g; S171, c; S191, a; S204, a; S206, c; I207, gg; D208, 324 776 1-368 1200, c; S201, a; S291, ta; I332, c 777 5-207 D1, 4; S141, r; D343, 126 779 4-360 D1, 3; S13, m; S15, c; S22, s; S24, m; S48, r; S66, s; S335, c; S345, rs; I360, ggg; D361, 119 780 1-472 1347, c; D473, 32 781 116-426 D1, 115; S219, m; S424, g; D427, 118 782 1-391 S386, k; D392, 64 783 1-453 D109, 1; S110, y; S125, y; I128, g; S132, k; 1453, ctctc 784 29-494 D1, 28; S72, r; D495, 93 785 99-461 D1, 98; S218, r; 1461, gaccgggg 786 2-465 D1, 1; S8, y; S388, s; I398, g; S400, t; S403, at; S417, g; D466, 24 787 28-271 D1, 27; S99, t; S230, c; S266, ga; S269, c; 1271, g; D272, 126 788 1-285 D280, 1; 1285, g; D286, 310 790 51-297 D1, 50; 1297, ggggg; D298, 539 791 113-327 D1, 10; S91, c; 192, a; D93, 258 794 9-431 D1, 16; S58, t; S217, t; I218, gggg; D219, 219 793 11-92 D1, 10; S91, c; 192, a; D93, 258 794 9-431 D1, 24; S315, g; D286, 310 795 30-341 D1, 29; I341, a; D342, 175 796 1-442 S17, w; S19, wr; D35, 1; S134, t; S264, n; S322, nr; S369, s; S420, s; S422, y; I442, tcctggg 797 1-420 S136, c; S150, c; I245, ccc; I420, ggagtg 798 25-316 D1, 24; S315, g; D317, 97 799 1-344 D345, 57 800 7-465 D1, 6; S59, k; S146, a; S186, km; i465, gttca 801 121-422 D1, 124; S315, p; D317, 97 799 1-344 D345, 57 800 7-465 D1, 6; S59, k; S146, a; S186, km; i465, gttca 801 121-422 D1, 145; S132, bn; I477, actac	766	1-320	S278, ag; S281, cagacc; S288, ta; S291, caag;
768			S296, c; S317, m; I320, cggg; D321, 306
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	803	15-467	D1, 14; S45, k; S65, t; S418, ys; D452, 1;
804 1-341 S42, t; S97, d; S326, gtg; S331, tgt; S336, a;			
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S221, k; I247, ccccaggg 1601 1-225 S109, bm; S195, m; I225, tgcacggg 1602 23-245 D1, 22; D138, 1; S139, s; S242, t; S244, g; I245, g; D246, 13 1603 1-303 S71, c; D277, 1; I303, ggagggg; D304, 38 1604 1-242 S47, w; S50, c; S81, h; S85, d; S91, k; S106, r; I242, tgtggg; D243, 50 1605 2-225 D1, 1; S20, k; S91, c; I225, ggg; D226, 132 1606 15-293 D1, 14; S156, g; S193, g; I200, t; I293,
1601 1-225 S109, bm; S195, m; I225, tgcacggg 1602 23-245 D1, 22; D138, 1; S139, s; S242, t; S244, g; I245, g; D246, 13 1603 1-303 S71, c; D277, 1; I303, ggagggg; D304, 38 1604 1-242 S47, w; S50, c; S81, h; S85, d; S91, k; S106, r; I242, tgtggg; D243, 50 1605 2-225 D1, 1; S20, k; S91, c; I225, ggg; D226, 132 1606 15-293 D1, 14; S156, g; S193, g; I200, t; I293,
1602 23-245 D1, 22; D138, 1; S139, s; S242, t; S244, g; I245, g; D246, 13 1603 1-303 S71, c; D277, 1; I303, ggagggg; D304, 38 1604 1-242 S47, w; S50, c; S81, h; S85, d; S91, k; S106, r; I242, tgtggg; D243, 50 1605 2-225 D1, 1; S20, k; S91, c; I225, ggg; D226, 132 1606 15-293 D1, 14; S156, g; S193, g; I200, t; I293,
I245, g; D246, 13 1603 1-303 S71, c; D277, 1; I303, ggagggg; D304, 38 1604 1-242 S47, w; S50, c; S81, h; S85, d; S91, k; S106, r; I242, tgtggg; D243, 50 1605 2-225 D1, 1; S20, k; S91, c; I225, ggg; D226, 132 1606 15-293 D1, 14; S156, g; S193, g; I200, t; I293,
1603 1-303 S71, c; D277, 1; I303, ggagggg; D304, 38 1604 1-242 S47, w; S50, c; S81, h; S85, d; S91, k; S106, r; I242, tgtggg; D243, 50 1605 2-225 D1, 1; S20, k; S91, c; I225, ggg; D226, 132 1606 15-293 D1, 14; S156, g; S193, g; I200, t; I293,
1604 1-242 S47, w; S50, c; S81, h; S85, d; S91, k; S106, r; I242, tgtggg; D243, 50 1605 2-225 D1, 1; S20, k; S91, c; I225, ggg; D226, 132 1606 15-293 D1, 14; S156, g; S193, g; I200, t; I293,
I242, tgtggg; D243, 50 1605 2-225 D1, 1; S20, k; S91, c; I225, ggg; D226, 132 1606 15-293 D1, 14; S156, g; S193, g; I200, t; I293,
1605 2-225 D1, 1; S20, k; S91, c; I225, ggg; D226, 132 1606 15-293 D1, 14; S156, g; S193, g; I200, t; I293,
1606 15-293 D1, 14; S156, g; S193, g; I200, t; I293,
acaaaggg
1607 1-361 S323, c; I361, cccca
1608 1-151 I151, taagggg; D152, 154
1609 1-242 S55, s; I135, a; S152, h; I242, cagtaggg
1610 1-196 I151, w; S190, k; I196, cctgtgg
1611 1-228 S115, k; S174, rk; I228, cgtttggg
1612 1-221 S108, v; I221, tgatcggg
1613 1-281 166, w; 1137, a; D282, 79
1614 1-171 S53, k; S76, k; I80, k; S81, kw; S86, r; S92, k;
S126, k; I171, gccgagg
1615 2-193 D1, 1; S67, c; I121, s; S122, mm; S126, g;
S130, r; S146, r; S156, gm; I193, cctca
1616 1-349 S251, ww; S259, rs; S275, k; I279, w; S285, y;
1 10000 1000 1001 1000 1041
S292, y; I320, m; I331, m; I338, w; I341, s;
I349, accceggg
I349, accceggg 1617
I349, accccggg 1617
I349, accceggg 1617 1-129 I118, t; D130, 26 1618 1-184 D9, 1; D185, 1 1619 1-169 I122, t; I169, gcccaggg 1620 1-187 S106, k; S118, m; S122, cg; S132, k; D188, 59
I349, accccggg 1617

EXAMPLE 16

Categorization of 5' ESTs and Consensus Contigated 5'ESTs

The nucleic acid sequences of the present invention (SEQ ID NOs. 24-811 and 1600-1622) were grouped based on their homology to known sequences as rollows. All sequences were compared to EMBL release 57 and daily releases available at the time of filing using BLASTN. All matches with a minimum of 25 nucleotides with 90% homology were retrieved and used to compute Tables IV and V.

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In some embodiments, 5'ESTs or consensus contigated 5'ESTs nucleic acid sequence do not match any known vertebrate sequence nor any publicly available EST sequence, thus being completely new.

In other embodiments, 5'ESTs or consensus contigated 5'ESTs match a known sequence. 5 Tables III and IV gives for each sequence of the invention in this category referred to by its sequence identification number in the first column, the positions of their preferred fragments in the second column entitled "Positions of preferred fragments." As used herein the term "polynucleotide described in Table III" refers to the all of the preferred polynucleotide fragments defined in Table III in this manner, and the term "polynucleotide described in Table IV" refers to the all of the preferred polynucleotides fragments 10 defined in Table IV in this manner. The present invention encompasses isolated, purified, or recombinant nucleic acids which consist of, consist essentially of, or comprise a contiguous span of at least 8, 10, 12, 15, 18, 20, 25, 35, 40, 50, 70, 80, 100, 250, or 500 nucleotides in length, to the extent that a contiguous span of these lengths is consistent with the lengths of the particular polynucleotide, of a polynucleotide described in Table III or Table IV, or a sequence complementary thereto, wherein said 15 polynucleotide described in Table III or Talbe IV is selected individually or in any combination from the polynucleotides described in Table III or Talbe IV. The present invention also encompasses isolated, purified, or recombinant nucleic acids which consist of or consist essentially of a polynucleotide described in Table III or Table IV, or a sequence complementary thereto, wherein said polynucleotide is selected individually or in any combination from the polynucleotides described in Table III or Table IV.

Table III

SEQ ID	Positions of preferred
NO	fragments
24	1-251
25	1-83
28	227-276
29	1-27
30	130-242, 283-315, 365-461
32	314-399
33	89-321
34	1-38
35	1-52, 171-222
36	1-30, 408-441
37	1-138
39	115-140
40	1-97
41	1-112
42	i-i77
46	1-38
48	376-400
51	400-466
54	1-259
55	189-320

	38
56	265-457
58	246-469
59	81-123, 418-444
60	1-348
61	78-123, 418-457
62	386-439
63	1-214
64	109-297
65	1-370
66	92-428
68	1-180
69	165-259
70	1-178
70	1-27
72	1-179
73	1-65, 107-192
	1-03, 107-192
75	
77	263-388
78	1-64
79	1-149
80	101-142, 302-380
82	1-192
83	1-398
85	1-290
86	1-118, 149-336
87	1-262
88	1-149
89	1-315
90	1-74
91	1-335, 364-423
92	1-316
93	338-508
94	179-321
95	219-402
96	26-315
97	348-460
98	1-230
99	391-467
101	214-336
102	1-289
103	1-383
104	1-211
105	1-36
106	1-126
107	1-49
108	294-336
109	1-128
	1-154
111	407-441
112	
113	1-80, 139-184
114	10-79
116	1-292
117	1-304

119	1-288
120	2-348
121	1-122
123	188-353
124	1-249
125	295-375
	1-244
128	
129	1-232
130	196-312
131	178-276
132	37-174
133	1-344
134	1-244
135	1-217
136	82-428
137	1-29, 103-155, 274-434
138	1-395
139	1-268
140	1-170
141	1-396
142	1-73, 227-357
143	1-159
144	1-433
145	61-116
146	1-71, 179-205
147	177-300
149	1-146
151	1-166
152	1-382 1-208
153	
154	121-251
155	1-147
157	1-115
158	1-175
159	1-44, 80-230
160	1-346
161	1-277
162	1-235
163	1-34
164	1-195
165	19-78, 175-217
166	1-209
- 167	1-65
168	128-218
169	49-245
170	179-280
171	1-103
172	1-218
	1-380
173	
174	1-139
175	1-122
176	1-300
177	1-466

	40
179	1-86
180	1-245
181	1-241
182	1-263
183	1-170
184	58-106, 399-443
185	1-427
186	1-365
187	1-260
188	1-172
189	1-150
190	161-271, 301-339
191	1-91
192	1-264
193	1-246
193	1-150
195	1-209
196	1-363
197	1-155
198	1-135
200	1-125
201	1-210
202	1-338
203	1-188
204	228-347
205	1-440
206	56-221
208	1-422
209	169-195
210	1-363
211	1-368
212	1-448
213	1-134
214	1-193
215	1-214
216	1-134
218	1-189
219	1-248
220	1-115
221	1-113
222	1-370
224	1-251
225	1-198
226	45-141
227	1-206
228	1-480
229	1-144
230	1-42, 281-351, 432-457
231	1-112
233	1-301
234	1-109
235	1-393
236	1-222

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237	1-154		
238	1-439		
239	112-137		
240	1-194		
241	1-44		
242	1-242		
244	1-324		
245	. 1-38, 217-280		
246	1-60		
247	77-359		
248	1-236		
249	1-342		
250	80-382		
251	1-303		
252	62-259		
253	1-165		
254	1-328		
255	1-320		
256	1-305		
257	1-181		
258	116-174		
259	1-265		
260	1-272		
261	1-62		
263	1-371		
266	1-274		
267	1-342		
268	364-427		
269	31-143		
270	1-79		
271	1-121		
272	229-292		
273	1-158		
274	1-113		
275	1-254		
276	1-333		
277	1-130		
278	1-184		
279	1-265		
280	1-188		
281	1-177		
282	1-336		
	1-294		
283			
284	1-171		
285	1-297		
288	1-42		
290	1-170		
292	20-155		
294	1-334		
295	1-375		
296	1-226		
297	1-232		
299	40-139		

	42
300	1-285
301	1-242
302	1-136
303	1-175
304	1-493
305	1-214
	89-458
306	
307	1-328
308	1-380
309	1-236
310	1-357
311	1-470
312	1-187
313	1-159
315	1-162
	1-404
316	
317	1-450
318	1-395
319	1-257
320	56-325
321	1-201
322	1-159
323	1-420
324	1-210
325	1-192
326	88-181
327	1-185
328	128-210
330	1-223
331	1-362
332	1-89
334	1-188
335	1-115
336	1-300
337	1-307
	1-123
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339	1-297
340	1-34
341	1-44
342	1-37
343	141-169
344	1-112
345	1-235, 266-349
346	1-191
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347	1-229
348	1-210
350	139-266
351	1 307
352	1-170
353	1-293
354	30-161, 192-331
355	1-93
356	1-178
	1+1/0

357 1-107 358 1-29, 168-209 359 1-298 360 1-193 362 1-360	
359 1-298 360 1-193	
359 1-298 360 1-193	
362 1-360	
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363 1-45, 100-212	
364 39-170, 202-242	
365 1-248	
366 1-351	
367 1-208	
368 228-446	
369 1-62	
370 1-132	
371 1-127	
372 1-196	
373 1-148	
374 1-126	
375 1-112	
376 1-146	
378 1-143	
379 1-261	
380 202-228	
382 1-151	
383 1-45	
385 1-55, 141-181 386 1-281	
387 1-111	
388 1-374	
389 1-192	
390 1-371	
392 1-303	
394 1-126	
395 1-329	
396 1-99	
397 1-316 398 1-251	
399 1-120 401 1-206	
403 1-311	
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406 1-206	
407 1-479	
408 1-289	
410 229-321	
413 1-158	
415 95-229	
416 1-265	
417 1-228	
418 1-225	
419 207-293	
420 1-194	

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423	1-420		
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425	1-276, 309-419		
426	1-232		
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428	1-96		
429	1-165		
431	1-58, 186-237, 327-354		
433	1-65		
434	1-83		
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436	405-447		
438	1-106		
439	45-105, 168-255, 284-447		
441	1-409		
442	1-320		
443	1-320		
444	1-284		
445	1-240		
446			
	1-149 1-360		
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448	1-123		
449	1-94		
450	1-302		
452	1-349		
453	1-270		
454	1-492		
455	17-105		
456	1-102		
457	1-108		
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459	1-311		
460	1-191		
461	312-420		
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464	1-142		
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468	1-41		
469	1-438		
470	1-131		
471	1-211		
472	1-150		
473	1-352		
474	1-141		
476	1-232		
478	1-201		
479	1-151		
480	1-104		
481	7-429		
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486	1-226		
488	1-296		
489	1-72, 323-377		
491	1-348		
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497	1-255		
499	1-174, 384-474		
500	1-50, 102-241		
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503	21-63, 356-420		
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512	1-140, 170-246, 276-420		
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514	1-303		
515	13-340		
516	1-263, 293-360		
518	1-245		
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520	62-182		
521	1-218		
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525	1-276		
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538	1-415		
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542	1-38, 73-390		
	1-221		
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545	1-376		

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553	1-232	
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564	184-352	
566	308-341	
567	1-218	
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569	1-142	
570	1-207	
571	1-373	
572	1-195	
573	1-352	
574	1-121	
575	1-222	
576	151-288	
577	1-264	
578	1-205	
580	1-171, 273-328	
581	1-356	
582	1-239	
583	1-144	
584	1-282	
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586	1-436	
588	1-380	
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590	1-178	
	1-66	
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597	31-83	
598	1-417	
599	1-329	
600	1-311	
601	1-61, 99-214	
602	1-154, 197-463	
603	135-269	
604	1-351	
605	1-195	
608	1-357	
609	1-201	
	1-201	

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612	1-176
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615	1-272
616	1-114
617	1-46
618	1-208
619	1-257
620	1-28
621	1-26
622	1-221
623	1-432
624	1-233
625	1-26
627	1-43
628	1-318
629	1-170
630	1-196
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634	1-134
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642	1-309
643	1-75, 162-213
644	107-211
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646	1-347
647	1-49, 81-143
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659	1-197
660	116-172
661	1-411
662	1-146
663	1-65
664	1-182
665	1-320
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668	1-122
670	1-160

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671	1-137
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674	1-263
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680	1-26
682	1-58, 269-328
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686	1-60
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689	132-221, 327-377
690	1-388
691	1-141, 171-408
692	1-141, 171-408
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695	1-455
698	1-58, 117-174
699	240-300
700	1-159
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	1-69
702	1-175
703	1-298
704	1-136
705	1-168
706	1-419
707	1-382
708	8-245, 296-384
709	1-149
710	1-167
711	1-35
712	1-80, 116-156, 206-241
713	33-376
714	1-304
715	1-242
717	1-145
718	1-350
720	1-257
721	1-360
722	1-191
724	1-139
726	1-207
727	99-164
728	1-321
730	156-372
731	1-109, 256-290
735	25-192
737	1-160
738	1-227
739	441-514
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742 217-280 743 10-275 747 1-179 749 2-31, 139-168 750 349-410 752 1-119 753 1-121 754 1-28 760 25-175 761 1-212 763 8-75 766 1-59, 102-248, 295-320 769 53-85 771 1-370 774 1-347 776 1-200 778 39-342 779 4-28 780 1-49, 407-472 781 116-426 782 1-59 783 1-53, 219-453 784 29-53, 219-263, 426-494 785 99-347, 386-461 786 2-28 788 1-279 789 1-58 790 226-268 792 129-218 796 5-86 797 1-34			
747 1-179 749 2-31, 139-168 750 349-410 752 1-119 753 1-121 754 1-28 760 25-175 761 1-212 763 8-75 766 1-59, 102-248, 295-320 769 53-85 771 1-370 774 1-347 776 1-200 778 39-342 779 4-28 780 1-49, 407-472 781 116-426 782 1-59 783 1-53, 219-453 784 29-53, 219-263, 426-494 785 99-347, 386-461 786 2-28 788 1-279 789 1-58 790 226-268 792 129-218 794 265-431 796 5-86 797 1-34 806 64-384	742	217-280	
749 2-31, 139-168 750 349-410 752 1-119 753 1-121 754 1-28 760 25-175 761 1-212 763 8-75 766 1-59, 102-248, 295-320 769 53-85 771 1-370 774 1-347 776 1-200 778 39-342 779 4-28 780 1-49, 407-472 781 116-426 782 1-59 783 1-53, 219-453 784 29-53, 219-263, 426-494 785 99-347, 386-461 786 2-28 788 1-279 789 1-58 790 226-268 792 129-218 794 265-431 796 5-86 797 1-34 806 64-384 807 135-301	743	10-275	
749 2-31, 139-168 750 349-410 752 1-119 753 1-121 754 1-28 760 25-175 761 1-212 763 8-75 766 1-59, 102-248, 295-320 769 53-85 771 1-370 774 1-347 776 1-200 778 39-342 779 4-28 780 1-49, 407-472 781 116-426 782 1-59 783 1-53, 219-453 784 29-53, 219-263, 426-494 785 99-347, 386-461 786 2-28 788 1-279 789 1-58 790 226-268 792 129-218 794 265-431 796 5-86 797 1-34 806 64-384 807 135-301	747	1-179	
750			
752 1-119 753 1-121 754 1-28 760 25-175 761 1-212 763 8-75 766 1-59, 102-248, 295-320 769 53-85 771 1-370 774 1-347 776 1-200 778 39-342 779 4-28 780 1-49, 407-472 781 116-426 782 1-59 783 1-53, 219-453 784 29-53, 219-263, 426-494 785 99-347, 386-461 786 2-28 788 1-279 789 1-58 790 226-268 792 129-218 794 265-431 796 5-86 797 1-34 806 64-384 807 135-301 808 2-314 810 6-39 1601<	750		
753 1-121 754 1-28 760 25-175 761 1-212 763 8-75 766 1-59, 102-248, 295-320 769 53-85 771 1-370 774 1-347 776 1-200 778 39-342 779 4-28 780 1-49, 407-472 781 116-426 782 1-59 783 1-53, 219-453 784 29-53, 219-263, 426-494 785 99-347, 386-461 786 2-28 788 1-279 789 1-58 790 226-268 792 129-218 794 265-431 796 5-86 797 1-344 802 46-477 806 64-384 807 135-301 808 2-314 810 6-39 160		1-119	
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792 129-218 794 265-431 796 5-86 797 1-34 799 1-344 802 46-477 806 64-384 807 135-301 808 2-314 810 6-39 1600 1-25 1601 1-225 1602 23-139 1603 1-294 1606 15-44 1607 1-361 1611 85-228 1612 1-221 1613 138-281 1614 65-171 1615 2-142 1616 1-46 1617 1-95 1620 1-187	789	1-58	
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796 5-86 797 1-34 799 1-344 802 46-477 806 64-384 807 135-301 808 2-314 810 6-39 1600 1-25 1601 1-225 1602 23-139 1603 1-294 1606 15-44 1607 1-361 1611 85-228 1612 1-221 1613 138-281 1614 65-171 1615 2-142 1616 1-46 1617 1-95 1620 1-187	792		
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806 64-384 807 135-301 808 2-314 810 6-39 1600 1-25 1601 1-225 1602 23-139 1603 1-294 1606 15-44 1607 1-361 1611 85-228 1612 1-221 1613 138-281 1614 65-17i 1615 2-142 1616 1-46 1617 1-95 1620 1-187	802	46-477	
807 135-301 808 2-314 810 6-39 1600 1-25 1601 1-225 1602 23-139 1603 1-294 1606 15-44 1607 1-361 1611 85-228 1612 1-221 1613 138-281 1614 65-171 1615 2-142 1616 1-46 1617 1-95 1620 1-187			
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1600 1-25 1601 1-225 1602 23-139 1603 1-294 1606 15-44 1607 1-361 1611 85-228 1612 1-221 1613 138-281 1614 65-171 1615 2-142 1616 1-46 1617 1-95 1620 1-187			
1601 1-225 1602 23-139 1603 1-294 1606 15-44 1607 1-361 1611 85-228 1612 1-221 1613 138-281 1614 65-171 1615 2-142 1616 1-46 1617 1-95 1620 1-187			
1602 23-139 1603 1-294 1606 15-44 1607 1-361 1611 85-228 1612 1-221 1613 138-281 1614 65-171 1615 2-142 1616 1-46 1617 1-95 1620 1-187			
1603 1-294 1606 15-44 1607 1-361 1611 85-228 1612 1-221 1613 138-281 1614 65-171 1615 2-142 1616 1-46 1617 1-95 1620 1-187		<u></u>	
1606 15-44 1607 1-361 1611 85-228 1612 1-221 1613 138-281 1614 65-171 1615 2-142 1616 1-46 1617 1-95 1620 1-187			
1607 1-361 1611 85-228 1612 1-221 1613 138-281 1614 65-17i 1615 2-142 1616 1-46 1617 1-95 1620 1-187			
1611 85-228 1612 1-221 1613 138-281 1614 65-171 1615 2-142 1616 1-46 1617 1-95 1620 1-187		<u> </u>	
1612 1-221 1613 138-281 1614 65-171 1615 2-142 1616 1-46 1617 1-95 1620 1-187		<u> </u>	
1613 138-281 1614 65-17i 1615 2-142 1616 1-46 1617 1-95 1620 1-187			
1614 65-171 1615 2-142 1616 1-46 1617 1-95 1620 1-187			
1615 2-142 1616 1-46 1617 1-95 1620 1-187			
1616 1-46 1617 1-95 1620 1-187			
1617 1-95 1620 1-187			
1620 1-187			
1621 1-136			
	1621	1-136	

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1622	32-280	. 311	-400	

Table IV

CEO ID NO	Desitions of Duofound	
SEQ ID NO	Positions of Preferred Fragments	
35	1-52	
41	1-115	
45	1-47	
46	1-33	
66	400-428	
82	83-149	
93	399-508 1-36	
105		
114	1-79	
120	1-386	
121	1-190	
124	1-249	
125	295-328	
139	1-81, 125-268	
159	1-139, 180-230	
165	1-78	
170	179-205, 248-280	
194	1-150	
213	1-158	
247	1-104, 155-183, 280-359	
269	31-143	
350	139-386	
368	228-446	
385	1-72, 143-193	
415	95-229	
435	1-386	
436	446-472	
441	1-361	
454	1-349	
455	1-105	
459	35-161, 200-311	
460	1-26, 56-140	
481	1-429	
489	1-84	
496	1-44, 84-215	
501	153-430	
502	1-91	
504	1-63	
	<u></u> _	

51			
505	1-68		
514	1-303		
515	237-351		
519	1-145		
526	231-366		
530	1-88		
535	1-55		
570	76-207		
576	168-218, 261-288		
588	1-331		
597	1-83		
627	1-43		
634	1-41		
641	1-55, 334-483		
672	1-34		
687	1-129		
708	1-245, 296-384		
710	1-26, 104-167		
722	1-191		
730	1-465		
731	1-43		
735	1-91		
737	1-160		
738	1-186		
739	1-48		
742	1-62, 99-248		
743	1-315, 412-459		
744	1-31		
747	1-63		
749	1-32		
750	1-38		
752	1-139		
753	1-193		
754	1-28		
759	1-38		
760	1-115		
763	1-62		
765	1-126		
769	1-85		
770	1-40		
771	1-148		
774	1-134		
	<u> </u>		
775	265-531		
776	71-203		
777	333-469		
778	144-468		
779	1-28		
780	1-49		
781	1-102		
782	1-59		
783	1-53		
784	1-220, 262-390		
785	1-339, 408-461		

786 1-28 789 1-58 791 1-126 792 1-31, 129-220 793 1-31 794 355-431 795 1-33 797 1-31 798 1-31 799 1-401 801 1-117 802 1-92 806 64-384 807 1-331
791 1-126 792 1-31, 129-220 793 1-31 794 355-431 795 1-33 797 1-31 798 1-31 799 1-401 801 1-117 802 1-92 806 64-384
792 1-31, 129-220 793 1-31 794 355-431 795 1-33 797 1-31 798 1-31 799 1-401 801 1-117 802 1-92 806 64-384
793 1-31 794 355-431 795 1-33 797 1-31 798 1-31 799 1-401 801 1-117 802 1-92 806 64-384
793 1-31 794 355-431 795 1-33 797 1-31 798 1-31 799 1-401 801 1-117 802 1-92 806 64-384
795 1-3.3 797 1-31 798 1-31 799 1-401 801 1-117 802 1-92 806 64-384
797 1-31 798 1-31 799 1-401 801 1-117 802 1-92 806 64-384
798 1-31 799 1-401 801 1-117 802 1-92 806 64-384
799 1-401 801 1-117 802 1-92 806 64-384
801 1-117 802 1-92 806 64-384
802 1-92 806 64-384
806 64-384
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807 1-331
808 1-351
810 1-39
1600 1-25
1603 1-341
1606 1-31
1607 1-361
1608 164-305
1611 85-228
1612 1-221
1613 112-360
1614 1-171
1615 94-193
1617 1-155
1620 1-246

III. Evaluation of Spatial and Temporal Expression of mRNAs Corresponding to the 5'ESTs, Consensus Contigated 5'ESTs, or EST-related nucleic acids

5

10

EXAMPLE 17

Expression Patterns of mRNAs From Which the 5'ESTs were obtained

Each of the SEQ ID NOs. 24-811 and 1600-1622 was also categorized based on the tissue from which its corresponding mRNA was obtained, as follows.

Table V shows the spatial distribution of each nucleic acid sequence of the invention (SEQ ID NOs. 24-811 and 1600-1622) referred to by its sequence identification number in the first column. In the second column entitled tissue distribution, the spatial distribution is represented by the number of individual 5'ESTs used to assemble the consensus contigated 5'ESTs for a given tissue. Each type of fissue listed in Table V is encoded by a letter. The correspondence between the letter code and the tissue 15 type is given in Table VI.

Table V

SEQ ID NO	Tissue Distribution
24	AA:1
25	S:1
26	P:1
27	W:1
28	P:1
29	S:1
30	P:1
31	P:1
32	P:1
33	P:1
34	AB:1
35	G:3; P:1; S:1; W:3; AA:4
36	P:1
37	S:1
38	Q:1
39	P:1
40	AB:1
41	B:1; C:3; F:1; G:1; H:4; S:2; T:8; W:1; Z:1; AA:3; AC:1; AD:3
42	A:1
43	N:2
44	P:1
45	C:2; K:1; O:1; S:5
46	K:1; S:2; AA:1
47	AA:1
48	C:1; O:1; P:8
49	P:1
50	P:1
51	P:1
52	S:1
53	AA:1
54	T:1
55	P:1
56_	P:1
57	P:1
58	P:1
59	P:7; T:2; Z:1
60	R:1
бi	C:1
62	P:1
63	F:1
64	AA:1
65	F:1

	54
66	P:4; T:2; Z:1
67	S:1
68	AA:1
69	P:1
70	P:1
71	S:1
72	W:1
73	G:1
74	P:1
75	N:1
76	P:1
77	S:1
78	U:1
79	B:1
80	P:1
81	AC:1
82	K:1; O:1
83	G:1
84	C:1; K:2; P:29; S:2; T:1; X:2; Y:1; AA:2
85	K:1
86	C:1
87	F:1
88	AB:1
89	H:1
90	M:1
91	B:1
92	K:1
93	AC:2
94	P:1
95	M:1
96	Z:2
97	K:1; P:11; S:1; X:1; AA:1
98	W:1
99	X:1
100	P:1
101	AB:1
102	F:1
103	AA:1
104	K:1
105	B:4; C:6; E:2; H:3; O:2; Q:1; S:3; AC:2
106	T:1
107	0:1
108	P:1
109	G:1
110	AA:1
111	T:1
112	P:1
113	F:1

	55
114	B:3; C:4; K:5; S:4; Y:1
115	U:1
116	W:1
117	T:1
118	T:2
119	T:1
120	H:3
121	AA:3
122	K:1
123	H:2
124	AA:2
125	B:1; G:1; J:3; T:13; Y:5; AA:5; AD:2
126	H:1; P:1
127	K:1
128	F:1
129	G:1
130	P:1
131	B:1
132	AA:1
133	W:1
134	P:1
135	K:1
136	B:1; C:1
137	B:1
138	H:1
139	AC:2
140	T:1
141	B:1
142	H:1
143	T:1
144	H:1
145	B:1
146	R:1
147	P:1
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156	R:1
157	W:1
158	T:1
159	C:1; AA:1
160	F:1
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198 O:1		
199 B:1		
200 AA:1		
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202 B:1		
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205 AA:1		AA:
206 Y:1		
207 Y:1		Y:1
208 AA:1		
209 G:1	209	G:1

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210	H:1
211	C:1
212	H:1
213	W:2
214	Y:1
215	AB:1
216	K:1
217	M:1
218	AD:1
219	A:1
220	AA:1
221	G:1
222	G:1
223	G:1; H:2; S:2; X:1
224	G:1
225	G:1
226	B:1
227	P:1
228	0:1
229	G:1
230	T:1
231	T:1
232	K:1
233	S:1
234	O:1
235	F:1
236	T:1
237	B:1
238	W:1
239	G:1
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241	A:1
242	W:1
243	P:1
244	H:1
245	D:1
246	C:1
247	B:2
248	P:1
249	F:1
250	AB:1
251	W:1
252	H:1
253	B:1
254	S:1
255	T:1
256	W:1
257	T:1

258	AA:2
259	P:1
260	W:1
	H:1
	K:1
263	K:1
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265	A:1
266	T:1 K:1
267	
268	H:1
269	T:2
270	T:1
271	T:1
272	B:1
273	Y:1
274	T:1
275	G:1
276	AA:1
277	T:1
278	AB:1
279	T:1
280	W:1
281	F:1
282	K:1
283	H:1
284	O:1 W:1
285	B:21; C:7; H:5; K:5; O:8; S:16; W:1; Y:3; Z:4; AA:2; AC:1
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287	K:2; P:12; W:1; AC:2 S:1
288	
289	K:2; P:8; W:1; AC:2
290	S:1 H:1
291	B:11; C:2; E:1; H:7; K:1; N:3; S:1; T:8; W:1; AA:28; AC:1
292	B:6; C:3; G:1; H:6; K:4; N:4; O:3; Q:2; S:5; T:1; U:1; V:2; Y:3; AA:1
293	B:0; C:5; G:1; H:0; K:4; N:4; O:5; Q:2; 5:5; 1:1; O:1; V:2; 1:5; AR:1
294	H:1
	AA:1
296	T:1
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300	H:1; S:1
301	II.1
302	W:1
303	W:1
304	H:1
305	G:1

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306	K:1
307	H:1
308	A:1
309	H:1
310	H:1
311	Y:1
312	G:1
313	H:1
314	K:1
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316	P:1
317	H:1
318	AA:1
319	H:1
320	0:1
321	Y:1
322	B:1
323	P:1
324	P:1
325	K:1
326	H:1
327	H:1
328	Q:1 · · ·
329	S:1
330	B:1
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338	AA:1
339	AA:1
340	G:1
341	C:1
342	K:1
343	S:1
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345	B:1
346	Y:1
347	G:1
348	F:1
349	AA:5
350	B:15; C:1; G:1; H:1; O:1; Q:2; S:1; X:1; Y:1
351	F:1
352	R:1
353	0:1
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354 H:1 355 W:1 356 F:1 357 T:1 358 S:1 359 X:1 360 T:1 361 K:1 362 K:1	
356 F:1 357 T:1 358 S:1 359 X:1 360 T:1 361 K:1	
357 T:1 358 S:1 359 X:1 360 T:1 361 K:1	
358 S:1 359 X:1 360 T:1 361 K:1	
359 X:1 360 T:1 361 K:1	
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361 K:1	
363 G:1	
364 K:1	
365 G:1	
366 AA:1	
367 F:1	
368 C:2; H:2; X:1	
369 E:1	
370 T:1	
371 H:1	
372 G:1	
373 AA:1	
374 G:1	
375 F.1	
376 F:1	
377 R:1	
378 AA:1	
379 AA:1	
380 C:1	
381 H:1	
382 T:1	
383 W:1	
384 S:1	
385 AA:2	
386 D:1	
387 O:1	
388 W:1	
389 F:1	
390 W:1	
390 W:1 391 K:1	
394 T:1	
395 H:1	
396 T:1	
397 T:1	
398 G:1	
399 C:1	
400 K:1	
401 B:1	

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402	H:1
403	B:1
404	B:1
405	H:1
406	AB:1
407	O:1
408	P:1
409	X:1
410	H:1
411	B:9; C:3; K:3; L:2; O:1; S:2; X:1; AA:1
412	G:1; S:2; V:2; W:1; Y:1; Z:1
413	W:1
414	G:1
415	B:3; C:3; F:1; G:2; H:4; J:1; K:1; O:1; P:3; S:1; V:1
416	I:1
417	F:1
418	F:1
419	F:1
420	AA:1
421	F:1
422	T:1
423	P:1
424	B:1
425	Y:1
426	W:1
427	AA:1
428	W:1
429	H:1
430	Y:1
431	J:1
432	AA:1
433	G:1
434	AA:1
435	B:3; H:1
436	B:9; G:4; H:8; K:2; O:2; W:1; Z:2; AA:2; AD:3
437	H:1; T:1
438	T:1
439	R:1
440	M:1
441	H:2
442	W:1
443	B:1
444	W:1
445	AB:1
446	F:1
447	AD:1
448	AB:1
449	N:1

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451	W:1
452	O:1
453	AA:1
454	D:28
455	W:1
456	T:1
457	G:1
458	W:1
459	Y:4
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461	P:2
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464	H:1
465	G:1
466	AC:1
467	R:1
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469	B:1
470	S:1
471	T:1
472	AA:1
473	W:1
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484	W:1
485	P:1
486	B:1
487	Y:1
488	H:1
489	P:1; Q:1; S:3
490	C:1
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492	H:1
493	B:!
494	H:1
495	G:1
496	N:2
497	B:1

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499	P:1	
500	G:1	
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502	B:4	
503	R:1	
504	B:5; H:2; W:2	
505	G:2; H:1	
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507	B:1	
508	W:1	
509	AB:1	
510	H:1	
511	N:1	
512	J:1	
513	AA:1	
514	T:2	
515	AA:5	
516	F:1	
517	C:1; O:1	
518	W:1	
519	T:4	
520	B:1	
521	H:1	
522	H:2; T:3	
523	H:1	
524	AA:1	
525	W:1	
526	C:2; E:1; J:1; R:3; S:4; AA:1	
527	H:1	
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529	P:1	
530	B:1; H:1	
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532	Y:1	
533	H:1	
534	T:1	
535	T:2	
536	B:1	
537	AD:1	
538	AA:1	
539	T:1	
540	F:1	
541	AD:1	
542	W:1	
543	W:1	
544	F:1	
545	T:1	

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550	B:1	
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552	B:1	
553	H:1	
554	P:1	
555	G:1	
556	H:1	
557	K:1	
558	B:1	
559	R:1	
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569	W:1	
570	B:2	
571	O:1	
572	T:1	
573	B:1	
574	T:1	
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576	B:3	
577	B:1	
578	X:1	
579	H:1	
580	AA:1	
581	AA:1	
582	AA:1 .	
583	AA:1	
584	AA:1	
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586	H:1	
587	H:1	
588	AA:3	
589	K:1	
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594	V:1
595	R:1
596	P:1
597	G:1; X:2; Z:1
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599	F:1
600	F:1
601	Y:1
602	F:1
603	W:1
604	H:1
605	G:1
606	C:2; H:1; S:3; W:2; AD:3
607	W:1
608	C:1
609	F:1
610	K:1
611	M:1
612	AD:1
613	H:1
614	T:1
615	H:1
616	F:1
617	T:1
618	G:1
619	G:1
620	B:1
621	W:1
622	W:1
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624	AA:1
625	G:1
626	M:1
627	C:2; T:2; W:1; Y:1
628	T:1
629	J:1
630	T:1
631	P:1
632	H:1
633	H:1
634	C:1; S:1; T:1; AD:1
635	J:1
636	G:1
637	W:1
638	AA:1
639	W:1
640	B:6; C:3; G:1; H:2; K:6; O:4; Q:1; R:2; S:1; T:3; Y:3; Z:2; AA:2; AC:2; AD:3

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643	P:1
644	AA:1
645	T:1
646	K:1
647	F:1
648	F:1
649	F:1
650	T:1
651	W:1
652	T:1
653	T:1
654	P:1
655	B:1; H:2; N:1; T:3; Y:1
656	B:1
657	T:1
658	R:1
659	K:1
660	W:1
661	AA:1
662	Y:1
663	W:1
664	G:1
665	S:1
666	Y:1
667	F:1
668	T:1
669	B:1
670	F:1
671	T:1
672	A:2; B:6; C:1; G:1; H:3; J:1; L:1; P:2; Q:1; S:4; T:1; V:3; W:2; Y:1;
	AA:3; AD:2
673	T:1
674	G:1
675	F:1
676	M:1
677	G:1
678	Y:1
679	D:1
680	P:1
681	D:1
682	AA:1
683	G:1
684	K:1
685	G:1
686	P:1
687	B:3; C:2; D:2; E:2; J:4; V:2; AC:6

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689	S:1
690	AA:1
691	H:1
692	AA:1
693	S:1
694	AB:1
695	T:1
696	H:1
697	B:4; E:1; F:1; P:1; T:2; Z:2
698	O:1
699	W:1
700	S:1
701	0:1
702	B:1
703	AB:1
704	H:1
705	B:1
706	H:1
707	G:1
708	F:1; H:1; K:1; W:2; AA:1
709	H:1
710	T:2
711	C:1
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713	Y:1
714	C:1
715	Y:1
716	Z:1
717	P:1
718	G:1
719	S:1
720	K:1
721	M:1
721	T:2
723	O:1; P:2; S:2
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725	T:1
726	N:1
727	T:1
728	T:1
729	C:2; H:2; K:2; V:1; AC:1
730	B:7; H:2; Y:1
731	B:5; W:3
732	B:1; C:2; G:2; S:2; AA:9
733	B:6; C:2; G:1; H:10; O:2; P:6; Q:1; S:2; W:4; AC:2
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735	C:1; O:2

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739	B:3; C:8; D:1; E:6; G:3; H:11; I:1; J:1; N:1; O:3; P:12; Q:3; S:2; T:2 W:1; AC:1; AD:8	
740	H:2; Y:1	
741	C:2; H:1	
742	B:12; C:1; G:1; H:4; K:2; O:2; S:4; T:2; Y:2	
743	AA:4	
744	B:1; G:1; H:6; T:1; W:1	
745	C:7; E:1; G:3; H:2; P:2; S:2; T:1; W:1; AD:2	
746	G:2; S:1	
747	T:2	
748	S:3	
749	H:1; O:2; S:2	
750	Y:1; AD:1	
751	B:8; G:2; H:2; I:1; Q:2; S:2; T:1; W:2	
752	T:3	
753	P:4	
754	B:1; H:2	
755	B:7; C:1; G:6; H:2; K:1; U:2; V:1; Z:1	
756	C:1; H:1; J:2; O:2; S:1; T:2; W:1; AA:1	
757	B:1; C:1; K:3; S:1; V:1; Y:1	
758	E:1; H:2; K:1; P:1; Q:1; AD:5	
759	B:6; C:1; Y:1	
760	B:4	
761	W:2	
762	B:3; C:7; H:9; N:1; S:1; T:1; Y:1; AA:1	
763	N:1; S:1; AA:5	
764	H:3	
765	B:3; G:1; W:1	
766	H:2	
767	C:1; AA:3	
768	B:2; C:6; H:9; N:1; S:1; T:1; Y:1; AA:1	
769	A:1; B:4; C:4; F:4; G:6; H:10; K:2; O:8; P:2; R:1; S:8; T:2; W:3; AA:2; AC:1	
770	A:2; P:16; X:1	
771	AA:3	
772	O:4	
773	B:1; C:1; W:1	
774	P:2; X:4	
775	B:18; C:6; H:5; K:3; O:7; S:10; W:1; Y:3; Z:2; AA:2; AC:1	
776	H:7	
777	B·26; C:8; H:5; K:4; O:10; S:17; W:1; Y:4; Z:4; AA:4; AC:2	
778	B:6	
779	B:3; C:1; G:1; H:2; K:1; Q:1; S:8; W:2; Y:9; AA:4	
780	B:3; C:1; F:1; P:1; W:1; AC:1	
781	I:2; N:1; P:1; R:3; AA:1	
782	B:2	

/ 04	H.1; F.1; 5:4; AD:1
785	T:2
786	D:1; AC:9
787	H:1; L:1; S:1
788	B:6; S:4
789	S:1; T:1
790	B:1; C:2; H:5; W:1; AD:1
791	B:3; C:2; D:3; E:2; J:4; V:3; AC:5
792	B:3; D:1; K:2; S:2; Y:1
793	B:2; G:2; AA:1
794	B:25; C:4; D:1; E:1; F:3; G:6; J:1; K:6; N:1; O:1; P:2; R:1; S:3; T:2;
	W:2; X:1; Y:1; Z:1; AA:1; AC:2; AD:1
795	B:4; C:1; E:2; H:4; J:1; L:1; O:4; S:1; V:1; Y:3; Z:1
796	H:5
797	B:2; E:1; N:2
798	B:1; G:1; H:6; T:1; W:1
799	H:2
800	H:2; I:2; AA:1
801	A:2; B:4; C:14; D:1; H:2; K:1; N:2; S:4; T:1; W:2; AA:20
802	AA:17
803	B:2; G:3; H:3; S:1; U:1; AC:1; AD:2
804	C:1; S:2; T:2; X:2; AA:1; AC:1
805	B:5; C:6; D:5; H:17; J:2; K:4; N:1; O:6; P:2; S:5; T:5; W:1; X:1; Z:2; AA:13; AC:3
806	B:2; C:3; D:3; H:6; J:2; K:1; N:1; O:3; P:1; S:2; T:4; W:1; X:1; Z:1; AA:5; AC:1
807	H:1; AC:4
808	R:13
809	B:3; W:4
810	B:16; S:1; Y:14
811	B:8; C:5; G:1; H:1; K:5; O:2; Q:2; R:2; S:2; T:3; Y:4; Z:2; AA:1; AC:1; AD:2
1600	T:4
1601	AA:3
1602	C:3; H:1
1603	H:2; AC:2
1604	B:7; C:1; E:1; H:1; P:2; R:3; S:2; T:2; Z:3; AA:2
1605	C:4; H:3; O:1
1606	A:3; B:13; C:14; D:2; E:10; F:3; G:19; H:32; K:11; O:5; P:2; R:3; S:16;
1000	T:4; W:2; Y:10; Z:8; AA:1; AC:3
1607	T:3
1608	B:3; P:2
1609	R:4
1610	B:4
1611	B:3; T:1
1612	T:2
1613	V:5
1614	D:3
1014	ID.J

1615	AA:10	
1616	B:4	
1617	T:2	
1618	K:2; S:8; AA:1	
. 1619	B:2	
1620	W:2	
1621	H:1; AB:1	
1622	H:2	

Table VI

Tissue code	Tissue type
Α	Bone Marrow
В	Brain
С	Cancerous prostate
D	Cerebellum
E	Colon
F	Dystrophic muscle
G	Fetal brain
Н	Fetal kidney
I	Fetal liver
J	Heart
K	Hypertrophic prostate
L	Kidney
M	Large intestine
N	Liver
O P	Lung
P	Lymph ganglia
Q	Lymphocytes
R	Muscle
S	Prostate
Т	Ovary
U	Pancreas
V	Placenta
W	Spinal cord
X	Spleen
Y	Substantia nigra
Z	Surrenals
AA	Testis
AB	Thyroid
AC	Umbilical cord
AD	Uterus

In addition to categorizing the 5' ESTs and consensus contigated 5' ESTs with respect to their tissue of origin, the spatial and temporal expression patterns of the mRNAs corresponding to the 5' ESTs and consensus contigated 5' ESTs, as well as their expression levels, may be determined as described in Example 18 below.

Characterization of the spatial and temporal expression patterns and expression levels of these mRNAs is useful for constructing expression vectors capable of producing a desired level of gene product in a desired spatial or temporal manner, as will be discussed in more detail below.

Furthermore, 5' ESTs and consensus contigated 5' ESTs whose corresponding mRNAs are 5 associated with disease states may also be identified. For example, a particular disease may result from the lack of expression, over expression, or under expression of a mRNA corresponding to a 5' EST or consensus contigated 5' EST. By comparing mRNA expression patterns and quantities in samples taken from healthy individuals with those from individuals suffering from a particular disease, 5' ESTs or consensus contigated 5' ESTs responsible for the disease may be identified.

It will be appreciated that the results of the above characterization procedures for 5' ESTs and consensus contigated 5' ESTs also apply to extended cDNAs (obtainable as described below) which contain sequences adjacent to the 5' ESTs and consensus contigated 5' ESTs. It will also be appreciated that if desired, characterization may be delayed until extended cDNAs have been obtained rather than characterizing the 5' ESTs or consensus contigated 5' ESTs themselves.

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EXAMPLE 18

Evaluation of Expression Levels and Patterns of mRNAs Corresponding to EST-Related Nucleic Acids

Expression levels and patterns of mRNAs corresponding to EST-related nucleic acids may be 20 analyzed by solution hybridization with long probes as described in International Patent Application No. WO 97/05277. Briefly, an EST-related nucleic acid, fragment of an EST-related nucleic acid, positional segment of an EST-related nucleic acid, or fragment of a positional segment of an EST-related nucleic acid corresponding to the gene encoding the mRNA to be characterized is inserted at a cloning site immediately downstream of a bacteriophage (T3, T7 or SP6) RNA polymerase promoter to produce 25 antisense RNA. Preferably, the EST-related nucleic acid, fragment of an EST-related nucleic acid, positional segment of an EST-related nucleic acid, or fragment of a positional segment of an EST-related nucleic acid is 100 or more nucleotides in length. The plasmid is linearized and transcribed in the presence of ribonucleotides comprising modified ribonucleotides (i.e. biotin-UTP and DIG-UTP). An excess of this doubly labeled RNA is hybridized in solution with mRNA isolated from cells or tissues of 30 interest. The hybridizations are performed under standard stringent conditions (40-50°C for 16 hours in an 80% formamide, 0.4 M NaCl buffer, pH 7-8). The unhybridized probe is removed by digestion with ribonucleases specific for single-stranded RNA (i.e. RNases CL3, T1, Phy M, U2 or A). The presence of the biotin-UTP modification enables capture of the hybrid on a microtitration plate coated with streptavidin. The presence of the DIG modification enables the hybrid to be detected and quantified by 35 ELISA using an anti-DIG antibody coupled to alkaline phosphatase.

The EST-related nucleic acid, fragment of an EST-related nucleic acid, positional segment of an EST-related nucleic acid, or fragment of a positional segment of an EST-related nucleic acid may also be 72

tagged with nucleotide sequences for the serial analysis of gene expression (SAGE) as disclosed in UK
Patent Application No. 2 305 241 A. In this method, cDNAs are prepared from a cell, tissue, organism
or other source of nucleic acid for which gene expression patterns must be determined. The resulting
cDNAs are separated into two pools. The cDNAs in each pool are cleaved with a first restriction

5 endonuclease, called an anchoring enzyme, having a recognition site which is likely to be present at least
once in most cDNAs. The fragments which contain the 5' or 3' most region of the cleaved cDNA are
isolated by binding to a capture medium such as streptavidin coated beads. A first oligonucleotide linker
having a first sequence for hybridization of an amplification primer and an internal restriction site for a
so called tagging endonuclease is ligated to the digested cDNAs in the first pool. Digestion with the

10 second endonuclease produces short tag fragments from the cDNAs.

A second oligonucleotide having a second sequence for hybridization of an amplification primer and an internal restriction site is ligated to the digested cDNAs in the second pool. The cDNA fragments in the second pool are also digested with the tagging endonuclease to generate short tag fragments derived from the cDNAs in the second pool. The tags resulting from digestion of the first and second pools with the anchoring enzyme and the tagging endonuclease are ligated to one another to produce so called ditags. In some embodiments, the ditags are concatamerized to produce ligation products containing from 2 to 200 ditags. The tag sequences are then determined and compared to the sequences of the EST-related nucleic acid, fragment of an EST-related nucleic acid, positional segment of an EST-related nucleic acid, or fragment of a positional segment of an EST-related nucleic acid to determine which 5' ESTs, consensus contigated 5' ESTs, or extended cDNAs are expressed in the cell, tissue, organism, or other source of nucleic acids from which the tags were derived. In this way, the expression pattern of the 5' ESTs, consensus contigated 5' ESTs, or extended cDNAs in the cell, tissue, organism, or other source of nucleic acids is obtained.

Quantitative analysis of gene expression may also be performed using arrays. As used herein,
the term array means a one dimensional, two dimensional, or multidimensional arrangement of ESTrelated nucleic acids, fragments of EST-related nucleic acids, positional segments EST-related nucleic
acids, or fragments of positional segments of EST-related nucleic acids. Preferably, the EST-related
nucleic acids, fragments of EST-related nucleic acids, positional segments EST-related nucleic acids, or
fragments of positional segments of EST-related nucleic acids are at least 10, 12, 15, 18, 20, 23, 25, 28,
30, 35, 40, or 50 nucleotides in length. More preferably, the EST-related nucleic acids, fragments of
EST-related nucleic acids, positional segments EST-related nucleic acids, or fragments of positional
segments of EST-related nucleic acids are at least 100 nucleotide long. More preferably, the fragments
are more than 100 nucleotides in length. In some embodiments, the EST-related nucleic acids,
fragments of EST-related nucleic acids, positional segments EST-related nucleic acids, or fragments of
positional segments of EST-related nucleic acids may be more than 500 nucleotides long.

For example, quantitative analysis of gene expression may be performed with EST-related nucleic acids, fragments of EST-related nucleic acids, positional segments EST-related nucleic acids, or

fragments of positional segments of EST-related nucleic acids in a complementary DNA microarray as described by Schena et al. (Science 270:467-470, 1995; Proc. Natl. Acad. Sci. U.S.A. 93:10614-10619, 1996). EST-related nucleic acids, fragments of EST-related nucleic acids, positional segments ESTrelated nucleic acids, or fragments of positional segments of EST-related nucleic acids are amplified by 5 PCR and arrayed from 96-well microtiter plates onto silylated microscope slides using high-speed robotics. Printed arrays are incubated in a humid chamber to allow rehydration of the array elements and rinsed, once in 0.2% SDS for 1 min, twice in water for 1 min and once for 5 min in sodium borohydride solution. The arrays are submerged in water for 2 min at 95°C, transferred into 0.2% SDS for 1 min, rinsed twice with water, air dried and stored in the dark at 25°C.

Cell or tissue mRNA is isolated or commercially obtained and probes are prepared by a single round of reverse transcription. Probes are hybridized to 1 cm² microarrays under a 14 x 14 mm glass coverslip for 6-12 hours at 60°C. Arrays are washed for 5 min at 25°C in low stringency wash buffer (1 x SSC/0.2% SDS), then for 10 min at room temperature in high stringency wash buffer (0.1 x SSC/0.2% SDS). Arrays are scanned in 0.1 x SSC using a fluorescence laser scanning device fitted with a custom 15 filter set. Accurate differential expression measurements are obtained by taking the average of the ratios of two independent hybridizations.

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Quantitative analysis of the expression of genes may also be performed with EST-related nucleic acids, fragments of EST-related nucleic acids, positional segments EST-related nucleic acids, or fragments of positional segments of EST-related nucleic acids in complementary DNA arrays as 20 described by Pietu et al. (Genome Research 6:492-503, 1996). The EST-related nucleic acids, fragments of EST-related nucleic acids, positional segments EST-related nucleic acids, or fragments of positional segments of EST-related nucleic acids thereof are PCR amplified and spotted on membranes. Then, mRNAs originating from various tissues or cells are labeled with radioactive nucleotides. After hybridization and washing in controlled conditions, the hybridized mRNAs are detected by phospho-25 imaging or autoradiography. Duplicate experiments are performed and a quantitative analysis of differentially expressed mRNAs is then performed.

Alternatively, expression analysis of the EST-related nucleic acids, fragments of EST-related nucleic acids, positional segments EST-related nucleic acids, or fragments of positional segments of EST-related nucleic acids can be done through high density nucleotide arrays as described by Lockhart 30 et al. (Nature Biotechnology 14: 1675-1680, 1996) and Sosnowsky et al. (Proc. Natl. Acad. Sci. 94:1119-1123, 1997). Oligonucleotides of 15-50 nucleotides corresponding to sequences of EST-related nucleic acids, fragments of EST-related nucleic acids, positional segments EST-related nucleic acids, or fragments of positional segments of EST related nucleic acids are synthesized directly on the chip (Lockhart et al., supra) or synthesized and then addressed to the chip (Sosnowsky et al., supra) 35 Preferably, the oligonucleotides are about 20 to 25 nucleotides in length.

cDNA probes labeled with an appropriate compound, such as biotin, digoxigenin or fluorescent dye, are synthesized from the appropriate mRNA population and then randomly fragmented to an

average size of 50 to 100 nucleotides. The said probes are then hybridized to the chip. After washing as described in Lockhart et al, supra and application of different electric fields (Sonowsky et al, supra.), the dyes or labeling compounds are detected and quantified. Duplicate hybridizations are performed. Comparative analysis of the intensity of the signal originating from cDNA probes on the same target 5 oligonucleotide in different cDNA samples indicates a differential expression of the mRNA corresponding to the 5' EST, consensus contigated 5' EST or extended cDNA from which the oligonucleotide sequence has been designed.

IV. Use of 5' ESTs to Clone Extended cDNAs and to Clone the Corresponding Genomic DNAs

Once 5' ESTs or consensus contigated 5' ESTs which include the 5' end of the corresponding mRNAs have been selected using the procedures described above, they can be utilized to isolate extended cDNAs which contain sequences adjacent to the 5' ESTs or consensus contigated 5' ESTs. The extended cDNAs may include the entire coding sequence of the protein encoded by the corresponding mRNA, including the authentic translation start site. If the extended cDNA encodes a 15 secreted protein, it may contain the signal sequence, and the sequence encoding the mature protein remaining after cleavage of the signal peptide.

Extended cDNAs which include the entire coding sequence of the protein encoded by the corresponding mRNA are referred to herein as "full-length cDNAs." Alternatively, the extended cDNAs may not include the entire coding sequence of the protein encoded by the corresponding mRNA, 20 although they do include sequences adjacent to the 5'ESTs or consensus contigated 5' ESTs. In some embodiments in which the extended cDNAs are derived from an mRNA encoding a secreted protein, the extended cDNAs may include only the sequence encoding the mature protein remaining after cleavage of the signal peptide, or only the sequence encoding the signal peptide.

Examples 19 and 20 below describe a general method for obtaining extended cDNAs using 5' 25 ESTs or consensus contigated 5' ESTs and nucleic acid homologous thereto. Example 21 below describes the cloning and sequencing of several extended cDNAs, including full-length cDNAs which include the authentic 5' end of the corresponding mRNA for several secreted proteins.

The methods of Examples 19 and 20 can also be used to obtain extended cDNAs which encode less than the entire coding sequence of proteins encoded by the genes corresponding to the 5' ESTs or 30 consensus contigated 5'ESTs. In some embodiments, the extended cDNAs isolated using these methods encode at least 5,10, 15, 20, 25, 30, 35, 40, 50, 75, 100, or 150 consecutive amino acids of one of the proteins encoded by the sequences of SEQ ID NOs. 24-811 and 1600-1622. In some embodiments, the extended cDNAs isolated using these methods encode at least 5, 10, 15, 20, 25, 30, 35, 40, 30, 75, 100, or 150 consecutive amino acids of one of the proteins encoded by the sequences of SFQ ID NOs. 24-

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General Method for Using 5' ESTs or Consensus Contigated 5'ESTs to Clone and Sequence Extended cDNAs which Include the Entire Coding Region and the Authentic 5'End of the Corresponding mRNA

The following general method may be used to quickly and efficiently isolate extended cDNAs including sequence adjacent to the sequences of the 5' ESTs or Consensus Contigated 5'ESTs used to obtain them. This method may be applied to obtain extended cDNAs for any 5' EST or consensus contigated 5' EST of the invention, including those 5' ESTs and consensus contigated 5' ESTs encoding secreted proteins. This method is illustrated in Figure 3.

1. Obtaining Extended cDNAs

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The method takes advantage of the known 5' sequence of the mRNA. A reverse transcription reaction is conducted on purified mRNA with a poly dT primer containing a nucleotide sequence at its 5' end allowing the addition of a known sequence at the end of the cDNA which corresponds to the 3' end of the mRNA. Such a primer and a commercially-available reverse transcriptase enzyme are added to a buffered mRNA sample yielding a reverse transcript anchored at the 3' polyA site of the RNAs. Nucleotide monomers are then added to complete the first strand synthesis.

After removal of the mRNA hybridized to the first cDNA strand by alkaline hydrolysis, the products of the alkaline hydrolysis and the residual poly dT primer can be eliminated with an exclusion column.

Subsequently, a pair of nested primers on each end is designed based on the known 5' sequence from the 5' EST or consensus contigated 5' EST and the known 3' end added by the poly dT primer used in the first strand synthesis. Software used to design primers are either based on GC content and melting temperatures of oligonucleotides, such as OSP (Illier and Green, *PCR Meth. Appl.* 1:124-128, 1991), or based on the octamer frequency disparity method (Griffais *et al.*, *Nucleic Acids Res.* 19: 3887-3891, 1991) such as PC-Rare (http:// bioinformatics.weizmann.ac.il/software/PC-Rare/doc/manuel.html). Preferably, the nested primers at the 5' end and the nested primers at the 3' end are separated from one another by four to nine bases. These primer sequences may be selected to have

A first PCR run is performed using the outer primer from each of the nested pairs. A second PCR run using the inner primer from each of the nested pairs is then performed on a small sample of the first PCR product. Thereafter, the primers and remaining nucleotide monomers are removed.

30 2. Sequencing Extended cDNAs or Fragments Thereof

melting temperatures and specificities suitable for use in PCR.

Due to the lack of position constraints on the design of 5' nested primers compatible for PCR use using the OSP software, amplicons of two types are obtained. Preferably, the second 5' primer is located upstream of the translation initiation codon thus yielding a nested PCR product containing the entire coding sequence. Such an extended cDNA may be used in a direct cloning procedure as described in section a below. However, in some cases, the second 5' primer is located downstream of the translation initiation codon, thereby yielding a PCR product containing only part of the ORF. Such incomplete PCR products are submitted to a modified procedure described in section b below.

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a) Nested PCR products containing complete ORFs

When the resulting nested PCR product contains the complete coding sequence, as predicted from the 5'EST or consensus contigated 5' EST sequence, it is directly cloned in an appropriate vector as described in section 3.

5 b) Nested PCR products containing incomplete ORFs

When the amplicon does not contain the complete coding sequence, intermediate steps are necessary to obtain both the complete coding sequence and a PCR product containing the full coding sequence. The complete coding sequence can be assembled from several partial sequences determined directly from different PCR products.

Once the full coding sequence has been completely determined, new primers compatible for PCR use are then designed to obtain amplicons containing the whole coding region. However, in such cases, 3' primers compatible for PCR use are located inside the 3' UTR of the corresponding mRNA, thus yielding amplicons which lack part of this region, i.e. the polyA tract and sometimes the polyadenylation signal, as illustrated in Figure 3. Such extended cDNAs are then cloned into an 15 appropriate vector as described in section 3.

c) Sequencing extended cDNAs

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Sequencing of extended cDNAs can be performed using a Die Terminator approach with the AmpliTaq DNA polymerase FS kit available from Perkin Elmer.

In order to sequence long PCR fragments, primer walking is performed using software such as 20 OSP to choose primers and automated computer software such as ASMG (Sutton et al., Genome Science Technol. 1: 9-19, 1995) to construct contigs of walking sequences including the initial 5' tag. Preferably, primer walking is performed until the sequences of full length cDNAs are obtained.

Completion of the sequencing of a given extended cDNA fragment may be assessed by comparing the sequence length to the size of the corresponding nested PCR product. When Northern 25 blot data are available, the size of the mRNA detected for a given PCR product may also be used to finally assess that the sequence is complete. Sequences which do not fulfill these criteria are discarded and will undergo a new isolation procedure.

3. Cloning Extended cDNAs

The PCR product containing the full coding sequence is then cloned in an appropriate vector. 30 For example, the extended cDNAs can be cloned into any expression vector known in the art, such as pED6dpc2 (DiscoverEase, Genetics Institute, Cambridge, MA).

Cloned PCR products are then entirely sequenced in order to obtain at least two sequences per clone. Preferably, the sequences are obtained from both sense and antisense strands according to --- the aforementioned procedure with the following modifications. First, both 5' and 3' ends of cloned 35 PCR products are sequenced in order to confirm the identity of the clone. Second, primer walking is performed if the full coding coding region has not been obtained yet. Contigation is then performed using primer walking sequences for cloned products as well as walking sequences that have already

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contigated for uncloned PCR products. The sequence is considered complete when the resulting contigs include the whole coding region as well as overlapping sequences with vector DNA on both ends. All the contigated sequences for each cloned amplicon are then used to obtain a consensus sequence.

5 4. Selection of Cloned Full length Sequences

a) Computer analysis of extended cDNAs

Following identification of contaminants and masking of repeats, structural features, e.g. polyA tail and polyadenylation signal, of the sequences of extended cDNAs are subsequently determined using methods known to those skilled in the art. For example, algorithm, parameters and criteria defined in Figure 10 may be used. Briefly, a polyA tail is defined as a homopolymeric stretch of at least 11 A with at most one alternative base within it. The polyA tail search is restricted to the last 20 nucleotides of the sequence and limited to stretches of 11 consecutive A's because sequencing reactions are often not readable after such a polyA stretch. To search for a polyadenylation signal, the polyA tail is clipped from the full-length sequence. The 50 nucleotides preceding the polyA tail are searched for the canonic polyadenylation AAUAAA signal allowing one mismatch to account for possible sequencing errors as well as known variation in the canonical sequence of the polyadenylation signal.

Functional features, e.g. ORFs and signal sequences, of the sequences of extended cDNAs are subsequently determined as follows. The 3 upper strand frames of extended cDNAs are searched for ORFs defined as the maximum length fragments beginning with a translation initiation codon and ending with a stop codon. ORFs encoding at least 80 amino acids are preferred. If extended cDNAs encoding secreted proteins are desired, each found ORF is then scanned for the presence of a signal peptide using the matrix method described in Example 13.

Sequences of extended cDNAs are then compared, on a nucleotidic or proteic basis, to public sequences available at the time of filing.

b) Selection of full-length cDNAs of interest

A negative selection may then be performed in order to eliminate unwanted cloned sequences resulting from either contaminants or PCR artifacts as follows. Sequences matching contaminant sequences such as vector DNA, tRNA, mtRNA, rRNA sequences are discarded as well as those encoding ORF sequences exhibiting extensive homology to repeats. Sequences obtained by direct cloning (section 1a) but lacking polyA tail may be discarded. Only ORFs ending either before the polyA tail (section 1a) or before the end of the cloned 3'UTR (section 1b) may be selected. If extended cDNAs encoding secreted proteins are desired, ORFs containing a signal peptide are considered. In addition, ORFs containing unlikely mature proteins such as mature proteins which size is less than 20 amino acids or less than 25% of the immature protein size may be eliminated.

Then, for each remaining full length cDNA containing several ORFs, a preselection of ORFs may be performed using the following criteria. The longest ORF is preferred. If extended cDNAs

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encoding secreted proteins are desired and if the ORF sizes are similar, the chosen ORF is the one which signal peptide has the highest score according to Von Heijne method.

Sequences of full length cDNA clones may then be compared pairwise after masking of the repeat sequences. Full-length cDNA sequences exhibiting extensive homology may be clustered in the 5 same class. Each cluster may then be subjected to a cluster analysis that detects sequences resulting from internal priming or from alternative splicing, identical sequences or sequences with several frameshifts. A selection may be operated between clones belonging to the same class in order to detect clones encoding homologous but distinct ORFs which may be both selected if they both contain sequences of interest.

Selection of full-length cDNA clones encoding sequences of interest may subsequently be performed using the following criteria. Structural parameters (initial tag, polyadenylation site and signal) are first checked. Then, homologies with known nucleic acids and proteins are examined in order to determine whether the clone sequence match a known nucleotide/protein sequence and, in the latter case, its covering rate and the date at which the sequence became public. If there is no extensive 15 match with sequences other than ESTs or genomic DNA, or if the clone sequence brings substantial new information, such as encoding a protein resulting from alternative splicing of an mRNA coding for an already known protein, the sequence is kept. Examples of such cloned full-length cDNAs containing sequences of interest are described in Example 21. Sequences resulting from chimera or double inserts or located on chromosome breaking points as assessed by homology to other sequences may be 20 discarded during this procedure.

Extended cDNAs prepared as described above may be subsequently engineered to obtain nucleic acids which include desired portions of the extended cDNA using conventional techniques such as subcloning, PCR, or in vitro oligonucleotide synthesis. For example, nucleic acids which include only the full coding sequences may be obtained using techniques known to those skilled in the art.

25 Alternatively, conventional techniques may be applied to obtain nucleic acids which contain only part of the coding sequences. In the case of nucleic acids encoding secreted proteins, nucleic acids containing only the coding sequence for the mature protein remaining after the signal peptide is cleaved off or nucleic acids which contain only the coding sequences for the signal peptides may be obtained.

Similarly, nucleic acids containing any other desired portion of the coding sequences for the 30 encoded protein may be obtained. For example, the nucleic acid may contain at least 10, 15, 18, 20, 25, 28, 30, 35, 40, 50, 75, 100, 150, 200, 300, 400 or 500 consecutive bases of an extended cDNA.

Once an extended cDNA has been obtained, it can be sequenced to determine the amino acid sequence it encodes. Once the encoded amino acid sequence has been determined, one can create and identify any of the many conceivable cDNAs that will-encode that protein by sirmly using the 35 degeneracy of the genetic code. For example, allelic variants or other homologous nucleic acids can be identified as described below. Alternatively, nucleic acids encoding the desired amino acid sequence can be synthesized in vitro.

In a preferred embodiment, the coding sequence may be selected using the known codon or codon pair preferences for the host organism in which the cDNA is to be expressed.

In addition to PCR based methods for obtaining cDNAs which include the authentic 5'end of the corresponding mRNA as well as the complete protein coding sequence of the corresponding mRNA, 5 traditional hybridization based methods may also be employed. These methods may also be used to obtain the genomic DNAs which encode the mRNAs from which the 5' ESTs or consensus contigated 5' ESTS were derived, mRNAs corresponding to the extended cDNAs, or nucleic acids which are homologous to extended cDNAs, 5' ESTs, or consensus contigated 5' ESTs. Example 19 below provides examples of such methods.

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EXAMPLE 20

Methods for Obtaining Extended cDNAs which Include the Entire Coding Region and the Authentic 5'End of the Corresponding mRNA or Nucleic Acids Homologous to Extended cDNAs, 5' ESTs or Consensus Contigated 5' ESTs

A full-length cDNA library can be made using the strategies described in Example 7. Alternatively, a cDNA library or genomic DNA library may be obtained from a commercial source or made using techniques familiar to those skilled in the art.

Such cDNA or genomic DNA libraries may be used to isolate extended cDNAs obtained from 5' ESTs or consensus contigated 5' ESTs or nucleic acids homologous to extended cDNAs, 5' ESTs, or 20 consensus contigated 5' ESTs as follows. The cDNA library or genomic DNA library is hybridized to a detectable probe. The detectable probe may comprise at least 10, 15, 18, 20, 25, 28, 30, 35, 40, 50, 75, 100, 150, 200, 300, 400 or 500 consecutive nucleotides of the 5' EST, consensus contigated 5' EST, or extended cDNA.

Techniques for identifying cDNA clones in a cDNA library which hybridize to a given probe 25 sequence are disclosed in Sambrook et al., Molecular Cloning: A Laboratory Manual 2d Ed., Cold Spring Harbor Laboratory Press, 1989. The same techniques may be used to isolate genomic DNAs. Briefly, cDNA or genomic DNA clones which hybridize to the detectable probe are identified and isolated for further manipulation as follows. The detectable probe described in the preceding paragraph is labeled with a detectable label such as a radioisotope or a fluorescent molecule. Techniques for 30 labeling the probe are well known and include phosphorylation with polynucleotide kinase, nick translation, in vitro transcription, and non radioactive techniques. The cDNAs or genomic DNAs in the library are transferred to a nitrocellulose or nylon filter and denatured. After blocking of non specific sites, the filter is incubated with the labeled probe for an amount of time sufficient to allow binding of the probe to cDNAs or genomic DNAs containing a sequence canable of hybridizing therein.

By varying the stringency of the hybridization conditions used to identify cDNAs or genomic DNAs which hybridize to the detectable probe, cDNAs or genomic DNAs having different levels of homology to the probe can be identified and isolated as described below.

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1. Identification of cDNA or Genomic DNA Sequences Having a High Degree of Homology to the Labeled Probe

To identify cDNAs or genomic DNAs having a high degree of homology to the probe sequence, the melting temperature of the probe may be calculated using the following formulas:

For probes between 14 and 70 nucleotides in length the melting temperature (Tm) is calculated using the formula: Tm=81.5+16.6(log (Na+))+0.41(fraction G+C)-(600/N) where N is the length of the probe.

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If the hybridization is carried out in a solution containing formamide, the melting temperature may be calculated using the equation Tm=81.5+16.6(log (Na+))+0.41(fraction G+C)-(0.63% 10 formamide)-(600/N) where N is the length of the probe.

Prehybridization may be carried out in 6X SSC, 5X Denhardt's reagent, 0.5% SDS, 100 µg denatured fragmented salmon sperm DNA or 6X SSC, 5X Denhardt's reagent, 0.5% SDS, 100 µg denatured fragmented salmon sperm DNA, 50% formamide. The formulas for SSC and Denhardt's solutions are listed in Sambrook et al., supra.

Hybridization is conducted by adding the detectable probe to the prehybridization solutions listed above. Where the probe comprises double stranded DNA, it is denatured before addition to the hybridization solution. The filter is contacted with the hybridization solution for a sufficient period of time to allow the probe to hybridize to extended cDNAs or genomic DNAs containing sequences complementary thereto or homologous thereto. For probes over 200 nucleotides in length, the 20 hybridization may be carried out at 15-25°C below the Tm. For shorter probes, such as oligonucleotide probes, the hybridization may be conducted at 15-25°C below the Tm. Preferably, for hybridizations in 6X SSC, the hybridization is conducted at approximately 68°C. Preferably, for hybridizations in 50% formamide containing solutions, the hybridization is conducted at approximately 42°C.

All of the foregoing hybridizations would be considered to be under "stringent" conditions.

Following hybridization, the filter is washed in 2X SSC, 0.1% SDS at room temperature for 15 minutes. The filter is then washed with 0.1X SSC, 0.5% SDS at room temperature for 30 minutes to 1 hour. Thereafter, the solution is washed at the hybridization temperature in 0.1X SSC, 0.5% SDS. A final wash is conducted in 0.1X SSC at room temperature.

cDNAs or genomic DNAs which have hybridized to the probe are identified by autoradiography 30 or other conventional techniques.

2. Obtaining cDNA or Genomic DNA Sequences Having Lower Degrees of Homology to the Labeled Probe

The above procedure may be modified to identify cDNAs or genored DNAs having decreasing -levels of homology to the probe sequence. For example, to obtain cDNAs or genomic DNAs of 35 decreasing homology to the detectable probe, less stringent conditions may be used. For example, the hybridization temperature may be decreased in increments of 5°C from 68°C to 42°C in a hybridization buffer having a sodium concentration of approximately 1M. Following hybridization, the filter may be

washed with 2X SSC, 0.5% SDS at the temperature of hybridization. These conditions are considered to be "moderate" conditions above 50°C and "low" conditions below 50°C.

Alternatively, the hybridization may be carried out in buffers, such as 6X SSC, containing formamide at a temperature of 42°C. In this case, the concentration of formamide in the hybridization 5 buffer may be reduced in 5% increments from 50% to 0% to identify clones having decreasing levels of homology to the probe. Following hybridization, the filter may be washed with 6X SSC, 0.5% SDS at 50°C. These conditions are considered to be "moderate" conditions above 25% formamide and "low" conditions below 25% formamide. cDNAs or genomic DNAs which have hybridized to the probe are identified by autoradiography.

3. Determination of the Degree of Homology between the Obtained cDNAs or Genomic DNAs and 5'ESTs, Consensus Contigated 5'ESTs, or Extended cDNAs or Between the Polypeptides Encoded by the Obtained cDNAs or Genomic DNAs and the Polypeptides Encoded by the 5'ESTs, Consensus Contigated 5'ESTs, or Extended cDNAs

To determine the level of homology between the hybridized cDNA or genomic DNA and the
5'EST, consensus contigated 5'EST or extended cDNA from which the probe was derived, the
nucleotide sequences of the hybridized nucleic acid and the 5'EST, consensus contigated 5'EST or
extended cDNA from which the probe was derived are compared. The sequences of the 5'EST,
consensus contigated 5'EST or extended cDNA from which the probe was derived and the sequences of
the cDNA or genomic DNA which hybridized to the detectable probe may be stored on a computer
readable medium as described below and compared to one another using any of a variety of algorithms
familiar to those skilled in the art, those described below.

To determine the level of homology between the polypeptide encoded by the hybridizing cDNA or genomic DNA and the polypeptide encoded by the 5'EST, consensus contigated 5'EST or extended cDNA from which the probe was derived, the polypeptide sequence encoded by the hybridized nucleic acid and the polypeptide sequence encoded by the 5'EST, consensus contigated 5'EST or extended cDNA from which the probe was derived are compared. The sequences of the polypeptide encoded by the 5'EST, consensus contigated 5'EST or extended cDNA from which the probe was derived and the polypeptide sequence encoded by the cDNA or genomic DNA which hybridized to the detectable probe may be stored on a computer readable medium as described below and compared to one another using any of a variety of algorithms familiar to those skilled in the art, those described below.

Protein and/or nucleic acid sequence homologies may be evaluated using any of the variety of sequence comparison algorithms and programs known in the art. Such algorithms and programs include, but are by no means limited to, TBLASTN, BLASTP, FASTA, TFASTA, and CLUSTALW (Pearson and Lipman, 1988, Proc. Natl. Acad. Sci. USA 85(8).2444-2448: Altschul et al., 1990, J. Mol. Biol. 215(3):403-410; Thompson et al., 1994, Nucleic Acids Res. 22(2):4673-4680; Higgins et al., 1996, Methods Enzymol. 266:383-402; Altschul et al., 1990, J. Mol. Biol. 215(3):403-410;

Altschul et al., 1993, Nature Genetics 3:266-272).

In a particularly preferred embodiment, protein and nucleic acid sequence homologies are evaluated using the Basic Local Alignment Search Tool ("BLAST") which is well known in the art (see, e.g., Karlin and Altschul, 1990, Proc. Natl. Acad. Sci. USA 87:2267-2268; Altschul et al., 1990, J. Mol. Biol. 215:403-410; Altschul et al., 1993, Nature Genetics 3:266-272; Altschul et al., 1997, Nuc. Acids Res. 25:3389-3402). In particular, five specific BLAST programs are used to perform the

- (1) BLASTP and BLAST3 compare an amino acid query sequence against a protein sequence database;
- (2) BLASTN compares a nucleotide query sequence against a nucleotide sequence database;

following task:

- (3) BLASTX compares the six-frame conceptual translation products of a query nucleotide sequence (both strands) against a protein sequence database;
- (4) TBLASTN compares a query protein sequence against a nucleotide sequence database translated in all six reading frames (both strands); and
- 15 (5) TBLASTX compares the six-frame translations of a nucleotide query sequence against the six-frame translations of a nucleotide sequence database.

The BLAST programs identify homologous sequences by identifying similar segments, which are referred to herein as "high-scoring segment pairs," between a query amino or nucleic acid sequence and a test sequence which is preferably obtained from a protein or nucleic acid sequence database. High-scoring segment pairs are preferably identified (i.e., aligned) by means of a scoring matrix, many of which are known in the art. Preferably, the scoring matrix used is the BLOSUM62 matrix (Gonnet et al., 1992, Science 256:1443-1445; Henikoff and Henikoff, 1993, Proteins 17:49-61). Less preferably, the PAM or PAM250 matrices may also be used (see, e.g., Schwartz and Dayhoff, eds., 1978, Matrices for Detecting Distance Relationships: Atlas of Protein Sequence and Structure, Washington: National Biomedical Research Foundation)

The BLAST programs evaluate the statistical significance of all high-scoring segment pairs identified, and preferably selects those segments which satisfy a user-specified threshold of significance, such as a user-specified percent homology. Preferably, the statistical significance of a high-scoring segment pair is evaluated using the statistical significance formula of Karlin (see, e.g., Karlin and Altschul, 1990, Proc. Natl. Acad. Sci. USA 87:2267-2268).

The parameters used with the above algorithms may be adapted depending on the sequence length and degree of homology studied. In some embodiments, the parameters may be the default parameters used by the algorithms in the absence of instructions from the user.

extended cDNA, 5'EST, or 5' consensus contigated 5'EST from which the probe was derived may be determined using the FASTDB algorithm described in Brutlag et al. Comp. App. Biosci. 6:237-245, 1990. In such analyses the parameters may be selected as follows: Matrix=Unitary, k-tuple=4,

Mismatch Penalty=1, Joining Penalty=30, Randomization Group Length=0, Cutoff Score=1, Gap Penalty=5, Gap Size Penalty=0.05, Window Size=500 or the length of the sequence which hybridizes to the probe, whichever is shorter. Because the FASTDB program does not consider 5' or 3' truncations when calculating homology levels, if the sequence which hybridizes to the probe is truncated relative to 5 the sequence of the extended cDNA, 5'EST, or consensus contigated 5'EST from which the probe was derived the homology level is manually adjusted by calculating the number of nucleotides of the extended cDNA, 5'EST, or consensus contigated 5' EST which are not matched or aligned with the hybridizing sequence, determining the percentage of total nucleotides of the hybridizing sequence which the non-matched or non-aligned nucleotides represent, and subtracting this percentage from the 10 homology level. For example, if the hybridizing sequence is 700 nucleotides in length and the extended cDNA, 5'EST, or consensus contigated 5' EST sequence is 1000 nucleotides in length wherein the first 300 bases at the 5' □end of the extended cDNA, 5'EST, or consensus contigated 5' EST are absent from the hybridizing sequence, and wherein the overlapping 700 nucleotides are identical, the homology level would be adjusted as follows. The non-matched, non-aligned 300 bases represent 30% of the length of 15 the extended cDNA, 5'EST, or consensus contigated 5' EST. If the overlapping 700 nucleotides are 100% identical, the adjusted homology level would be 100-30=70% homology. It should be noted that the preceding adjustments are only made when the non-matched or non-aligned nucleotides are at the 5'or 3'ends. No adjustments are made if the non-matched or non-aligned sequences are internal or under any other conditions.

For example, using the above methods, nucleic acids having at least 95% nucleic acid homology, at least 96% nucleic acid homology, at least 97% nucleic acid homology, at least 98% nucleic acid homology, at least 99% nucleic acid homology, or more than 99% nucleic acid homology to the extended cDNA, 5'EST, or consensus contigated 5' EST from which the probe was derived may be obtained and identified. Such nucleic acids may be allelic variants or related nucleic acids from other 25 species. Similarly, by using progressively less stringent hybridization conditions one can obtain and identify nucleic acids having at least 90%, at least 85%, at least 80% or at least 75% homology to the extended cDNA, 5'EST, or consensus contigated 5' EST from which the probe was derived.

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Using the above methods and algorithms such as FASTA with parameters depending on the sequence length and degree of homology studied, for example the default parameters used by the 30 algorithms in the absence of instructions from the user, one can obtain nucleic acids encoding proteins having at least 99%, at least 98%, at least 97%, at least 96%, at least 95%, at least 90%, at least 85%, at least 80% or at least 75% homology to the protein encoded by the extended cDNA, 5'EST, or consensus contigated 5' EST from which the probe was derived. In some embodiments, the homology levels can he determined using the "default" opening penalty and the "default" gap penalty, and a scoring matrix 35 such as PAM 250 (a standard scoring matrix; see Dayhoff et al., in: Atlas of Protein Sequence and Structure, Vol. 5, Supp. 3 (1978)).

Alternatively, the level of polypeptide homology may be determined using the FASTDB algorithm described by Brutlag et al. Comp. App. Biosci. 6:237-245, 1990. In such analyses the parameters may be selected as follows: Matrix=PAM 0, k-tuple=2, Mismatch Penalty=1, Joining Penalty=20, Randomization Group Length=0, Cutoff Score=1, Window Size=Sequence Length, Gap 5 Penalty=5, Gap Size Penalty=0.05, Window Size=500 or the length of the homologous sequence, whichever is shorter. If the homologous amino acid sequence is shorter than the amino acid sequence encoded by the extended cDNA, 5'EST, or consensus contigated 5' EST as a result of an N terminal and/or C terminal deletion the results may be manually corrected as follows. First, the number of amino acid residues of the amino acid sequence encoded by the extended cDNA, 5'EST, or consensus 10 contigated 5' EST which are not matched or aligned with the homologous sequence is determined. Then, the percentage of the length of the sequence encoded by the extended cDNA, 5'EST, or consensus contigated 5' EST which the non-matched or non-aligned amino acids represent is calculated. This percentage is subtracted from the homology level. For example wherein the amino acid sequence encoded by the extended cDNA, 5'EST, or consensus contigated 5' EST is 100 amino acids in length 15 and the length of the homologous sequence is 80 amino acids and wherein the amino acid sequence encoded by the extended cDNA or 5'EST is truncated at the N terminal end with respect to the homologous sequence, the homology level is calculated as follows. In the preceding scenario there are 20 non-matched, non-aligned amino acids in the sequence encoded by the extended cDNA, 5'EST, or consensus contigated 5' EST. This represents 20% of the length of the amino acid sequence encoded by 20 the extended cDNA, 5'EST, or consensus contigated 5' EST. If the remaining amino acids are 1005 identical between the two sequences, the homology level would be 100%-20%=80% homology. No adjustments are made if the non-matched or non-aligned sequences are internal or under any other conditions.

In addition to the above described methods, other protocols are available to obtain extended cDNAs using 5' ESTs or consensus contigated 5'ESTs as outlined in the following paragraphs.

Extended cDNAs may be prepared by obtaining mRNA from the tissue, cell, or organism of interest using mRNA preparation procedures utilizing polyA selection procedures or other techniques known to those skilled in the art. A first primer capable of hybridizing to the polyA tail of the mRNA is hybridized to the mRNA and a reverse transcription reaction is performed to generate a first cDNA strand.

The first cDNA strand is hybridized to a second primer containing at least 10 consecutive nucleotides of the sequences of SEQ ID NOs 24-811 and 1600-1622. Preferably, the primer comprises at least 10, 12, 15, 17, 18, 20, 23, 25, or 28 consecutive nucleotides from the sequences of SEQ ID NOs 24-811 and 1600-1622. In some embodiments, the primer comprises more than 30 nucleotides from the sequences of SEQ ID NOs 24-811 and 1600-1622. If it is desired to obtain extended cDNAs containing the full protein coding sequence, including the authentic translation initiation site, the second primer used contains sequences located upstream of the translation initiation site. The second primer is

extended to generate a second cDNA strand complementary to the first cDNA strand. Alternatively, RT-PCR may be performed as described above using primers from both ends of the cDNA to be obtained.

Extended cDNAs containing 5' fragments of the mRNA may be prepared by hybridizing an mRNA comprising the sequences of SEQ ID NOs. 24-811 and 1600-1622 with a primer comprising a complementary to a fragment of an EST-related nucleic acid hybridizing the primer to the mRNAs, and reverse transcribing the hybridized primer to make a first cDNA strand from the mRNAs. Preferably, the primer comprises at least 10, 12, 15, 17, 18, 20, 23, 25, or 28 consecutive nucleotides of the sequences complementary to SEQ ID NOs. 24-811 and 1600-1622.

Thereafter, a second cDNA strand complementary to the first cDNA strand is synthesized. The second cDNA strand may be made by hybridizing a primer complementary to sequences in the first cDNA strand to the first cDNA strand and extending the primer to generate the second cDNA strand.

The double stranded extended cDNAs made using the methods described above are isolated and cloned. The extended cDNAs may be cloned into vectors such as plasmids or viral vectors capable of replicating in an appropriate host cell. For example, the host cell may be a bacterial, mammalian, avian, or insect cell.

Techniques for isolating mRNA, reverse transcribing a primer hybridized to mRNA to generate a first cDNA strand, extending a primer to make a second cDNA strand complementary to the first cDNA strand, isolating the double stranded cDNA and cloning the double stranded cDNA are well known to those skilled in the art and are described in Current Protocols in Molecular Biology, John Wiley & Sons, Inc. 1997 and Sambrook et al., Molecular Cloning: A Laboratory Manual, Second Edition, Cold Spring Harbor Laboratory Press, 1989.

Alternatively, other procedures may be used for obtaining full-length cDNAs or extended cDNAs. In one approach, full-length or extended cDNAs are prepared from mRNA and cloned into double stranded phagemids as follows. The cDNA library in the double stranded phagemids is then rendered single stranded by treatment with an endonuclease, such as the Gene II product of the phage F1 and an exonuclease (Chang *et al.*, Gene 127:95-8, 1993). A biotinylated oligonucleotide comprising the sequence of a fragment of an EST-related nucleic acid is hybridized to the single stranded phagemids. Preferably, the fragment comprises at least 10, 12, 15, 17, 18, 20, 23, 25, or 28 consecutive nucleotides of the sequences of SEQ ID NOs. 24-811 and 1600-1622.

Hybrids between the biotinylated oligonucleotide and phagemids are isolated by incubating the hybrids with streptavidin coated paramagnetic beads and retrieving the beads with a magnet (Fry et al., Biotechniques, 13: 124-131, 1992). Thereafter, the resulting phagemids are released from the beads and converted into double stranded DNA using a primer specific for the 5' EST or consensus contigated 5'EST-sequence used to design the biotinylated oligonucleotide. Alternatively, protocols such as the Gene Trapper kit (Gibco BRL) may be used. The resulting double stranded DNA is transformed into bacteria. Extended cDNAs or full length cDNAs containing the 5' EST or consensus contigated 5'EST

sequence are identified by colony PCR or colony hybridization.

Using any of the above described methods in section III, a plurality of extended cDNAs containing full-length protein coding sequences or portions of the protein coding sequences may be provided as cDNA libraries for subsequent evaluation of the encoded proteins or use in diagnostic assays as described below.

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EXAMPLE 21

Full Length cDNAs

The procedures described in Example 19 and 20 were used to obtain extended cDNAs or full length cDNAs derived from 5' ESTs in a variety of tissues. The following list provides a few examples 10 of cDNAs obtained by these means.

Using this procedure, the full length cDNA of SEQ ID NO:1 (internal identification number 58-34-2-E7-FL2) was obtained. This cDNA encodes the signal peptide MWWFQQGLSFLPSALVIWTSA (SEQ ID NO:2) having a von Heijne score of 5.5.

Using this approach, the full length cDNA of SEQ ID NO:3 (internal identification number 48-15 19-3-G1-FL1) was obtained. This cDNA encodes the signal peptide MKKVLLLITAILAVAVG (SEQ ID NO: 4) having a von Heijne score of 8.2.

The full length cDNA of SEQ ID NO:5 (internal identification number 58-35-2-F10-FL2) was also obtained using this procedure. This cDNA encodes a signal peptide LWLLFFLVTAIHA (SEQ ID NO:6) having a von Heijne score of 10.7.

Furthermore, the polypeptides encoded by the extended or full-length cDNAs may be screened for the presence of known structural or functional motifs or for the presence of signatures, small amino acid sequences which are well conserved amongst the members of a protein family. The results obtained for the polypeptides encoded by a few full-length cDNAs derived from 5'ESTs that were screened for the presence of known protein signatures and motifs using the Proscan software from the GCG 25 package and the Prosite 15.0 database are provided below.

The protein of SEQ ID NO: 8 encoded by the full-length cDNA SEQ ID NO: 7 (internal designation 78-8-3-E6-CL0_1C) and expressed in adult prostate belong to the phosphatidylethanolamine-binding protein from which it exhibits the characteristic PROSITE signature from positions 90 to 112. Proteins from this widespread family, from nematodes to fly, 30 yeast, rodent and primate species, bind hydrophobic ligands such as phospholipids and nucleotides. They are mostly expressed in brain and in testis and are thought to play a role in cell growth and/or maturation, in regulation of the sperm maturation, motility and in membrane remodeling. They may act either through signal transduction or through oxidereduction reactions (for a review see -- Schoentgen and Jollès, FEBS Letters, 369-:22-26 (1995)). Taken together, these data suggest that the 35 protein of SEQ ID NO: 8 may play a role in cell growth, maturation and in membrane remodeling and/or may be related to male fertility. Thus, these protein may be useful in diagnosing and/or treating cancer, neurodegenerative diseases, and/or disorders related to male fertility and sterility.

The protein of SEQ ID No. 10 encoded by the full-length cDNA SEQ ID No. 9 (internal designation 108-013-5-O-H9-FLC) shows homologies with a family of lysophospholipases conserved among eukaryotes (yeast, rabbit, rodents and human). In addition, some members of this family exhibit a calcium-independent phospholipase A2 activity (Portilla et al, J. Am. Soc. Nephro., 9:1178-1186 (1998)). All members of this family exhibit the active site consensus GXSXG motif of carboxylesterases that is also found in the protein of SEQ ID No. 10 (position 54 to 58). In addition, this protein may be a membrane protein with one transmembrane domain as predicted by the software TopPred II (Claros and von Heijne, CABIOS applic. Notes, 10:685-686 (1994)). Taken together, these data suggest that the protein of SEQ ID NO:10 may play a role in fatty acid metabolism, probably as a phospholipase. Thus, this protein or part therein, may be useful in diagnosing and/or treating several disorders including, but not limited to, cancer, diabetes, and neurodegenerative disorders such as Parkinson's and Alzheimer's diseases. It may also be useful in modulating inflammatory responses to infectious agents and/or to suppress graft rejection.

The protein of SEQ ID NO: 12 encoded by the full-length cDNA SEQ ID NO: 11 (internal 15 designation 108-004-5-0-D10-FLC) shows remote homology to a subfamily of beta4galactosyltransferases widely conserved in animals (human, rodents, cow and chicken). Such enzymes, usually type II membrane proteins located in the endoplasmic reticulum or in the Golgi apparatus, catalyzes the biosynthesis of glycoproteins, glycolipid glycans and lactose. Their characteristic features defined as those of subfamily A in Breton et al, J. Biochem., 123:1000-1009 20 (1998) are pretty well conserved in the protein of SEQ ID NO: 12, especially the region I containing the DVD motif (positions 163-165) thought to be involved either in UDP binding or in the catalytic process itself. In addition, the protein of SEQ ID NO: 12 has the typical structure of a type II protein. Indeed, it contains a short 28-amino-acid-long N-terminal tail, a transmembrane segment from positions 29 to 49 and a large 278-amino-acid-long C-terminal tail as predicted by the software 25 TopPred II (Claros and von Heijne, CABIOS applic. Notes, 10:685-686 (1994)). Taken together, these data suggest that the protein of SEQ ID NO: 12 may play a role in the biosynthesis of polysaccharides, and of the carbohydrate moieties of glycoproteins and glycolipids and/or in cell-cell recognition. Thus, this protein may be useful in diagnosing and/or treating several types of disorders including, but not limited to, cancer, atherosclerosis, cardiovascular disorders, autoimmune disorders 30 and rheumatic diseases including rheumatoid arthritis.

The protein of SEQ ID NO: 14 encoded by the full-length cDNA SEQ ID NO: 13 (internal designation 108-009-5-0-A2-FLC) shows extensive homology to the bZIP family of transcription factors, and especially to the human luman protein (Lu et al., Mol. Cell. Biol., 17:5117-5126 (1997))). The match include the whole bZIP domain composed of a basic DNA-binding domain and of a leucine zipper allowing protein dimerization. The basic domain is conserved in the protein of SEQ ID NO: 14 as shown by the characteristic PROSITE signature (positions 224-237) except for a conservative substitution of a glutamic acid with an aspartic acid in position 233. The typical

PROSITE signature for leucine zipper is also present (positions 259 to 280). Taken together, these data suggest that the protein of SEQ ID NO: 14 may bind to DNA, hence regulating gene expression as a transcription factor. Thus, this protein may be useful in diagnosing and/or treating several types of disorders including, but not limited to, cancer.

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Bacterial clones containing plasmids containing the full length cDNAs described above are presently stored in the inventor's laboratories under the internal identification numbers provided above. The inserts may be recovered from the deposited materials by growing an aliquot of the appropriate bacterial clone in the appropriate medium. The plasmid DNA can then be isolated using plasmid isolation procedures familiar to those skilled in the art such as alkaline lysis minipreps or large scale 10 alkaline lysis plasmid isolation procedures. If desired the plasmid DNA may be further enriched by centrifugation on a cesium chloride gradient, size exclusion chromatography, or anion exchange chromatography. The plasmid DNA obtained using these procedures may then be manipulated using standard cloning techniques familiar to those skilled in the art. Alternatively, a PCR can be done with primers designed at both ends of the insertion. The PCR product which corresponds to the cDNA insert 15 can then be manipulated using standard cloning techniques familiar to those skilled in the art.

V. Expression of Proteins or Polypeptides Encoded by EST-related nucleic acids or Fragments thereof

EST-related nucleic acids, fragments of EST-related nucleic acids, positional segments of EST-20 related nucleic acids, and fragments of positional segments of EST-related nucleic acids may be used to express the polypeptides which they encode. In particular, they may be used to express EST-related polypeptides, fragments of EST-related polypeptides, positional segments of EST-related polypeptides, or fragments of positional segments of EST-related polypeptides. In some embodiments, the ESTrelated nucleic acids, positional segments of EST-related nucleic acids, and fragments of positional 25 segments of EST-related nucleic acids may be used to express the full polypeptide (i.e. the signal peptide and the mature polypeptide) of a secreted protein, the mature protein (i.e. the polypeptide generated after cleavage of the signal peptide), or the signal peptide of a secreted protein. If desired, nucleic acids encoding the signal peptide may be used to facilitate secretion of the expressed protein. It will be appreciated that a plurality of EST-related nucleic acids, fragments of EST-related nucleic acids, 30 positional segments of EST-related nucleic acids, or fragments of positional segments of EST-related nucleic acids may be simultaneously cloned into expression vectors to create an expression library for analysis of the encoded proteins as described below.

To express their enc ded proteins, the EST-related nucleic acids, fragments of EST-related nucleic acids, positional segments of EST-related nucleic acids, or fragments of positional segments of EST-related nucleic acids are cloned into a suitable expression vector. In some instances, nucleic acids encoding EST-related polypeptides, fragments of EST-related polypeptides, positional segments of EST-related polypeptides or fragments of positional segments of EST-related polypeptides may be cloned into a suitable expression vector.

In some embodiments, the nucleic acids inserted into the expression vector may comprise the coding sequence of a sequence selected from the group consisting of SEQ ID NOs. 24-811. In other embodiments, the nucleic acids inserted into the expression vector may comprise may comprise the full coding sequence (*i.e.* the nucleotides encoding the signal peptide and the mature polypeptide) of one of SEQ ID Nos. 766-792. In some embodiments, the nucleic acid inserted into the expression vector may comprise the nucleotides of one of the sequences of SEQ ID Nos. 766-792 which encode the mature polypeptide (*i.e.* the nucleotides encoding the polypeptide generated after cleavage of the signal peptide). In further embodiments, the nucleic acids inserted into the expression vector may comprise the nucleotides of 24-728 and 766-792 which encode the signal peptide to facilitate secretion of the expressed protein. The nucleic acids inserted into the expression vectors may also contain sequences upstream of the sequences encoding the signal peptide, such as sequences which regulate expression levels or sequences which confer tissue specific expression.

The nucleic acid inserted into the expression vector may encode a polypeptide comprising the one of the sequences of SEQ ID Nos. 812-1599. In some embodiments, the nucleic acid inserted into the expression vector may encode the full polypeptide sequence (i.e. the signal peptide and the mature polypeptide) included in one of SEQ ID Nos. 1554-1580. In other embodiments, the nucleic acid inserted into the expression vector may encode the mature polypeptide (i.e. the polypeptide generated after cleavage of the signal peptide) included in one of the sequences of SEQ ID Nos. 1554-1580. In further embodiments, the nucleic acids inserted into the expression vector may encode the signal peptide included in one of the sequences of 812-1516 and 1554-1580.

The nucleic acid encoding the protein or polypeptide to be expressed is operably linked to a promoter in an expression vector using conventional cloning technology. The expression vector may be any of the mammalian, yeast, insect or bacterial expression systems known in the art. Commercially available vectors and expression systems are available from a variety of suppliers including Genetics Institute (Cambridge, MA), Stratagene (La Jolla, California), Promega (Madison, Wisconsin), and Invitrogen (San Diego, California). If desired, to enhance expression and facilitate proper protein folding, the codon context and codon pairing of the sequence may be optimized for the particular expression organism in which the expression vector is introduced, as explained by Hatfield, et al., U.S.

35 Patent No. 5,082,767.

The following is provided as one exemplary method to express the proteins encoded by the nucleic acids described above. In some instances the nucleic acid encoding the protein or polypeptide to

be expressed includes a methionine initiation codon and a polyA signal. If the nucleic acid encoding the polypeptide to be expressed lacks a methionine to serve as the initiation site, an initiating methionine can be introduced next to the first codon of the nucleic acid using conventional techniques. Similarly, if the nucleic acid encoding the protein or polypeptide to be expressed lacks a polyA signal, this sequence can 5 be added to the construct by, for example, splicing out the polyA signal from pSG5 (Stratagene) using BglI and SalI restriction endonuclease enzymes and incorporating it into the mammalian expression vector pXT1 (Stratagene). pXT1 contains the LTRs and a portion of the gag gene from Moloney Murine Leukemia Virus. The position of the LTRs in the construct allow efficient stable transfection. The vector includes the Herpes Simplex thymidine kinase promoter and the selectable neomycin gene. 10 The nucleic acid encoding the polypeptide to be expressed is obtained by PCR from the bacterial vector using oligonucleotide primers complementary to the nucleic acid encoding the protein or polypeptide to be expressed and containing restriction endonuclease sequences for Pst I incorporated into the 5'primer and BglII at the 5' end of 3' primer, taking care to ensure that the nucleic acid encoding the protein or polypeptide to be expressed is correctly positioned with respect to the poly A signal. The purified 15 fragment obtained from the resulting PCR reaction is digested with PstI, blunt ended with an exonuclease, digested with Bgl II, purified and ligated to pXT1, now containing a poly A signal and digested with BglⅡ.

The ligated product is transfected into mouse NIH 3T3 cells using Lipofectin (Life Technologies, Inc., Grand Island, New York) under conditions outlined in the product specification.

20 Positive transfectants are selected after growing the transfected cells in 600 µg/ml G418 (Sigma, St. Louis, Missouri).

Alternatively, the nucleic acid encoding the protein or polypeptide to be expressed may be cloned into pED6dpc2. The resulting pED6dpc2 constructs may be transfected into a suitable host cell, such as COS 1 cells. Methotrexate resistant cells are selected and expanded. The expressed protein or polypeptide may be isolated, purified, or enriched as described above.

To confirm expression of the desired protein or polypeptide, the proteins or polypeptides produced by cells containing a vector with a nucleic acid insert encoding the protein or polypeptide are compared to those lacking such an insert. The expressed proteins are detected using techniques familiar to those skilled in the art such as Coomassie blue or silver staining or using antibodies against the protein or polypeptide encoded by the nucleic acid insert. Antibodies capable of specifically recognizing the protein of interest may be generated using synthetic 15-mer peptides having a sequence encoded by the appropriate nucleic acid. The synthetic peptides are injected into mice to generate antibody to the polypeptide encoded by the nucleic acid.

If the proteins or polypeptides encoded by the nucleic acid inserts are secreted, medium

prepared from the host cells or organisms containing an expression vector which contains a nucleic acid insert encoding the desired protein or polypeptide is compared to medium prepared from the control cells or organism. The presence of a band in medium from the cells containing the nucleic acid insert which

is absent from preparations from the control cells indicates that the protein or polypeptide encoded by the nucleic acid insert is being expressed and secreted. Generally, the band corresponding to the protein encoded by the nucleic acid insert will have a mobility near that expected based on the number of amino acids in the open reading frame of the nucleic acid insert. However, the band may have a mobility different than that expected as a result of modifications such as glycosylation, ubiquitination, or enzymatic cleavage.

Alternatively, if the protein expressed from the above expression vectors does not contain sequences directing its secretion, the proteins expressed from host cells containing an expression vector with an insert encoding a secreted protein or portion thereof can be compared to the proteins expressed in control host cells containing the expression vector without an insert. The presence of a band in samples from cells containing the expression vector with an insert which is absent in samples from cells containing the expression vector without an insert indicates that the desired protein or portion thereof is being expressed. Generally, the band will have the mobility expected for the secreted protein or portion thereof. However, the band may have a mobility different than that expected as a result of modifications such as glycosylation, ubiquitination, or enzymatic cleavage.

The expressed protein or polypeptide may be purified, isolated or enriched using a variety of methods. In some methods, the protein or polypeptide may be secreted into the culture medium via a native signal peptide or a heterologous signal peptide operably linked thereto. In some methods, the protein or polypeptide may be linked to a heterologous polypeptide which facilitates its isolation, purification, or enrichment such as a nickel binding polypeptide. The protein or polypeptide may also be obtained by gel electrophoresis, ion exchange chromatography, size chromatography, hplc, salt precipitation, immunoprecipitation, a combination of any of the preceding methods, or any of the isolation, purification, or enrichment techniques familiar to those skilled in the art.

The protein encoded by the nucleic acid insert may also be purified using standard
immunochromatography techniques using immunoaffinity chromatography with antibodies directed against the encoded protein or polypeptide as described in more detail below. If antibody production is not possible, the nucleic acid insert encoding the desired protein or polypeptide may be incorporated into expression vectors designed for use in purification schemes employing chimeric polypeptides. In such strategies, the coding sequence of the nucleic acid insert is ligated in frame with the gene encoding the other half of the chimera. The other half of the chimera may be β-globin or a nickel binding polypeptide. A chromatography matrix having antibody to β-globin or nickel attached thereto is then used to purify the chimeric protein. Protease cleavage sites may be engineered between the β-globin gene or the nickel binding polypeptide and the extended cDNA or portion thereof. Thus, the two polypeptides of the chimera may be separated from one another by protease digestion.

One useful expression vector for generating β -globin chimerics is pSG5 (Stratagene), which encodes rabbit β -globin. Intron II of the rabbit β -globin gene facilitates splicing of the expressed transcript, and the polyadenylation signal incorporated into the construct increases the level of

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expression. These techniques as described are well known to those skilled in the art of molecular biology. Standard methods are published in methods texts such as Davis et al., (Basic Methods in Molecular Biology, L.G. Davis, M.D. Dibner, and J.F. Battey, ed., Elsevier Press, NY, 1986) and many of the methods are available from Stratagene, Life Technologies, Inc., or Promega. Polypeptide may 5 additionally be produced from the construct using in vitro translation systems such as the In vitro ExpressTM Translation Kit (Stratagene).

Following expression and purification of the proteins or polypeptides encoded by the nucleic acid inserts, the purified proteins may be tested for the ability to bind to the surface of various cell types as described in Example 23 below. It will be appreciated that a plurality of proteins expressed from 10 these nucleic acid inserts may be included in a panel of proteins to be simultaneously evaluated for the activities specifically described below, as well as other biological roles for which assays for determining activity are available.

EXAMPLE 23

Analysis of Secreted Proteins to Determine Whether they Bind to the Cell Surface

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The EST-related nucleic acids, fragments of EST-related nucleic acids, positional segments of EST-related nucleic acids, fragments of positional segments of EST-related nucleic acids, nucleic acids encoding the EST-related polypeptides, nucleic acids encoding fragments of the EST-related polypeptides, nucleic acids encoding positional segments of EST-related polypeptides, or nucleic acids encoding fragments of positional segments of EST-related polypeptides are cloned into expression 20 vectors such as those described in Example 22. The encoded proteins or polypeptides are purified, isolated, or enriched as described above. Following purification, isolation, or enrichment, the proteins or polypeptides are labeled using techniques known to those skilled in the art. The labeled proteins or polypeptides are incubated with cells or cell lines derived from a variety of organs or tissues to allow the proteins to bind to any receptor present on the cell surface. Following the incubation, the cells are 25 washed to remove non-specifically bound proteins or polypeptides. The specifically bound labeled proteins or polypeptides are detected by autoradiography. Alternatively, unlabeled proteins or polypeptides may be incubated with the cells and detected with antibodies having a detectable label, such as a fluorescent molecule, attached thereto.

Specificity of cell surface binding may be analyzed by conducting a competition analysis in 30 which various amounts of unlabeled protein or polypeptide are incubated along with the labeled protein or polypeptide. The amount of labeled protein or polypeptide bound to the cell surface decreases as the amount of competitive unlabeled protein or polypeptide increases. As a control, various amounts of an unlabeled protein or polypeptide unrelated to the labeled protein or polypeptide is included in some. binding reactions. The amount of labeled protein or polypeptide bound to the cell surface does not 35 decrease in binding reactions containing increasing amounts of unrelated unlabeled protein, indicating that the protein or polypeptide encoded by the nucleic acid binds specifically to the cell surface.

As discussed above, human proteins have been shown to have a number of important physiological effects and, consequently, represent a valuable therapeutic resource. The human proteins or polypeptides made as described above may be evaluated to determine their physiological activities as described below.

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EXAMPLE 24

Assaying the Expressed Proteins or Polypeptides for Cytokine, Cell Proliferation or Cell Differentiation Activity

As discussed above, some human proteins act as cytokines or may affect cellular proliferation or 10 differentiation. Many protein factors discovered to date, including all known cytokines, have exhibited activity in one or more factor dependent cell proliferation assays, and hence the assays serve as a convenient confirmation of cytokine activity. The activity of a protein or polypeptide of the present invention is evidenced by any one of a number of routine factor dependent cell proliferation assays for cell lines including, without limitation, 32D, DA2, DA1G, T10, B9, B9/11, BaF3, MC9/G, M⁺ (preB 15 M⁺), 2E8, RB5, DA1, 123, T1165, HT2, CTLL2, TF-1, Mo7c and CMK. The proteins or polypeptides prepared as described above may be evaluated for their ability to regulate T cell or thymocyte proliferation in assays such as those described above or in the following references: Current Protocols in Immunology, Ed. by J.E. Coligan et al., Greene Publishing Associates and Wiley-Interscience, Takai et al. J. Immunol. 137:3494-3500, 1986., Bertagnolli et al. J. Immunol. 145:1706-1712, 1990., 20 Bertagnolli et al., Cellular Immunology 133:327-341, 1991. Bertagnolli, et al. J. Immunol. 149:3778-

3783, 1992; Bowman et al., J. Immunol. 152:1756-1761, 1994.

In addition, numerous assays for cytokine production and/or the proliferation of spleen cells, lymph node cells and thymocytes are known. These include the techniques disclosed in Current Protocols in Immunology. J.E. Coligan et al. Eds., 1:3.12.1-3.12.14, John Wiley and Sons, Toronto.

25 1994; and Schreiber, R.D. In Current Protocols in Immunology., supra 1: 6.8.1-6.8.8.

The proteins or polypeptides prepared as described above may also be assayed for the ability to regulate the proliferation and differentiation of hematopoietic or lymphopoietic cells. Many assays for such activity are familiar to those skilled in the art, including the assays in the following references: Bottomly et al., In Current Protocols in Immunology., supra. 1: 6.3.1-6.3.12,; deVries et al., J. Exp. 30 Med. 173:1205-1211, 1991; Moreau et al., Nature 36:690-692, 1988; Greenberger et al., Proc. Natl. Acad. Sci. U.S.A. 80:2931-2938, 1983; Nordan, R., In Current Protocols in Immunology., supra. 1: 6.6.1-6.6.5; Smith et al., Proc. Natl. Acad. Sci. U.S.A. 83:1857-1861, 1986; Bennett et al in Current Protocols in Immunology supra 1: 6.15.1; Ciarletta et al In Current Protocols in Immunology. supra 1. ó.13.1.

The proteins or polypeptides prepared as described above may also be assayed for their ability to regulate T-cell responses to antigens. Many assays for such activity are familiar to those skilled in the art, including the assays described in the following references: Chapter 3 (In vitro Assays for Mouse

Lymphocyte Function), Chapter 6 (Cytokines and Their Cellular Receptors) and Chapter 7. (Immunologic Studies in Humans) in Current Protocols in Immunology supra; Weinberger et al., Proc. Natl. Acad. Sci. USA 77:6091-6095, 1980; Weinberger et al., Eur. J. Immun. 11:405-411, 1981; Takai et al., J. Immunol. 137:3494-3500, 1986; Takai et al., J. Immunol. 140:508-512, 1988.

Those proteins or polypeptides which exhibit cytokine, cell proliferation, or cell differentiation activity may then be formulated as pharmaceuticals and used to treat clinical conditions in which induction of cell proliferation or differentiation is beneficial. Alternatively, as described in more detail below, nucleic acids encoding these proteins or polypeptides or nucleic acids regulating the expression of these proteins or polypeptides may be introduced into appropriate host cells to increase or decrease the 10 expression of the proteins or polypeptides as desired.

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EXAMPLE 25

Assaying the Expressed Proteins or Polypeptides for Activity as Immune System Regulators

The proteins or polypeptides prepared as described above may also be evaluated for their effects 15 as immune regulators. For example, the proteins or polypeptides may be evaluated for their activity to influence thymocyte or splenocyte cytotoxicity. Numerous assays for such activity are familiar to those skilled in the art including the assays described in the following references: Chapter 3 (In vitro Assays for Mouse Lymphocyte Function 3.1-3.19) and Chapter 7 (Immunologic studies in Humans) in Current 20 Protocols in Immunology, J.E. Coligan et al. Eds, Greene Publishing Associates and Wiley-Interscience; Herrmann et al., Proc. Natl. Acad. Sci. USA 78:2488-2492, 1981; Herrmann et al., J. Immunol. 128:1968-1974, 1982; Handa et al., J. Immunol. 135:1564-1572, 1985; Takai et al., J. Immunol. 137:3494-3500, 1986; Takai et al., J. Immunol. 140:508-512, 1988; Bowman et al., J. Virology 61:1992-1998; Bertagnolli et al. Cell. Immunol. 133:327-341, 1991; Brown et al., J. Immunol. 153:3079-3092, 25 1994.

The proteins or polypeptides prepared as described above may also be evaluated for their effects on T-cell dependent immunoglobulin responses and isotype switching. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references: Maliszewski, J. Immunol. 144:3028-3033, 1990; Mond et al. in Current Protocols in Immunology, 1: 30 3.8.1-3.8.16, supra.

The proteins or polypeptides prepared as described above may also be evaluated for their effect on immune effector cells, including their effect on Th1 cells and cytotoxic lymphocytes. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references: Chapter 3 (In vitro Assays for Mouse Lymphocyte Function 3.1-3.19) and Chapter 35 7 (Immunologic Studies in Humans) in Current Protocols in Immunology, supra; Takai et al., J. Immunol. 137:3494-3500, 1986; Takai et al., J. Immunol. 140:508-512, 1988; Bertagnolli et al., J. Immunol. 149:3778-3783, 1992.

The proteins or polypeptides prepared as described above may also be evaluated for their effect on dendritic cell mediated activation of naive T-cells. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references: Guery et al., J. Immunol. 134:536-544, 1995; Inaba et al., J. Exp. Med. 173:549-559, 1991; Macatonia et al., J. 5 Immunol, 154:5071-5079, 1995; Porgador et al J. Exp. Med 182:255-260, 1995; Nair et al., J. Virol. 67:4062-4069, 1993; Huang et al., Science 264:961-965, 1994; Macatonia et al J. Exp. Med 169:1255-1264, 1989; Bhardwaj et al., Journal of Clinical Investigation 94:797-807, 1994; and Inaba et al., J. Exp. Med 172:631-640, 1990.

The proteins or polypeptides prepared as described above may also be evaluated for their 10 influence on the lifetime of lymphocytes. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references: Darzynkiewicz et al., Cytometry 13:795-808, 1992; Gorczyca et al., Leukemia 7:659-670, 1993; Gorczyca et al., Cancer Res. 53:1945-1951, 1993; Itoh et al., Cell 66:233-243, 1991; Zacharchuk, J. Immunol. 145:4037-4045, 1990; Zamai et al., Cytometry 14:891-897, 1993; Gorczyca et al., Int. J. Oncol. 1:639-648, 1992.

The proteins or polypeptides prepared as described above may also be evaluated for their influence on early steps of T-cell commitment and development. Numerous assays for such activity are familiar to those skilled in the art, including without limitation the assays disclosed in the following references: Antica et al., Blood 84:111-117, 1994; Fine et al., Cell. Immunol. 155:111-122, 1994; Galy et al., Blood 85:2770-2778, 1995; Toki et al., Proc. Nat. Acad Sci. USA 88:7548-7551, 1991.

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Those proteins or polypeptides which exhibit activity as immune system regulators activity may then be formulated as pharmaceuticals and used to treat clinical conditions in which regulation of immune activity is beneficial. For example, the protein or polypeptide may be useful in the treatment of various immune deficiencies and disorders (including severe combined immunodeficiency), e.g., in regulating (up or down) growth and proliferation of T and/or B lymphocytes, as well as effecting the 25 cytolytic activity of NK cells and other cell populations. These immune deficiencies may be genetic or be caused by viral (e.g., HIV) as well as bacterial or fungal infections, or may result from autoimmune disorders. More specifically, infectious diseases caused by viral, bacterial, fungal or other infection may be treatable using the protein or polypeptide including infections by HTV, hepatitis viruses, herpesviruses, mycobacteria, Leishmania spp., plamodium. and various fungal infections such as 30 candidiasis. Of course, in this regard, a protein or polypeptide may also be useful where a boost to the immune system generally may be desirable, i.e., in the treatment of cancer.

Alternatively, the proteins or polypeptides prepared as described above may be used in treatment of autoimmune disorders including, for example, connective fissue disease, multiple sclerosis, systemic lupus erythematosus, rheumato d arthritis, autoimmune pulmonary inflammation, Guillain-35 Barre syndrome, autoimmune thyroiditis, insulin dependent diabetes mellitis, myasthenia gravis, graftversus-host disease and autoimmune inflammatory eye disease. Such a protein or polypeptide may also to be useful in the treatment of allergic reactions and conditions, such as asthma (particularly allergic

asthma) or other respiratory problems. Other conditions, in which immune suppression is desired (including, for example, organ transplantation), may also be treatable using the protein or polypeptide.

Using the proteins or polypeptides of the invention it may also be possible to regulate immune responses either up or down. Down regulation may involve inhibiting or blocking an immune response already in progress or may involve preventing the induction of an immune response. The functions of activated T-cells may be inhibited by suppressing T cell responses or by inducing specific tolerance in T cells, or both. Immunosuppression of T cell responses is generally an active non-antigen-specific process which requires continuous exposure of the T cells to the suppressive agent. Tolerance, which involves inducing non-responsiveness or anergy in T cells, is distinguishable from immunosuppression in that it is generally antigen-specific and persists after the end of exposure to the tolerizing agent. Operationally, tolerance can be demonstrated by the lack of a T cell response upon reexposure to specific antigen in the absence of the tolerizing agent.

Down regulating or preventing one or more antigen functions (including without limitation B lymphocyte antigen functions, such as, for example, B7 costimulation), e.g., preventing high level 15 lymphokine synthesis by activated T cells, will be useful in situations of tissue, skin and organ transplantation and in graft-versus-host disease (GVHD). For example, blockage of T cell function should result in reduced tissue destruction in tissue transplantation. Typically, in tissue transplants, rejection of the transplant is initiated through its recognition as foreign by T cells, followed by an immune reaction that destroys the transplant. The administration of a molecule which inhibits or blocks 20 interaction of a B7 lymphocyte antigen with its natural ligand(s) on immune cells (such as a soluble, monomeric form of a peptide having B7-2 activity alone or in conjunction with a monomeric form of a peptide having an activity of another B lymphocyte antigen (e.g., B7-1, B7-3) or blocking antibody), prior to transplantation, can lead to the binding of the molecule to the natural ligand(s) on the immune cells without transmitting the corresponding costimulatory signal. Blocking B lymphocyte antigen 25 function in this matter prevents cytokine synthesis by immune cells, such as T cells, and thus acts as an immunosuppressant. Moreover, the lack of costimulation may also be sufficient to anergize the T cells, thereby inducing tolerance in a subject. Induction of long-term tolerance by B lymphocyte antigenblocking reagents may avoid the necessity of repeated administration of these blocking reagents. To achieve sufficient immunosuppression or tolerance in a subject, it may also be necessary to block the 30 function of a combination of B lymphocyte antigens.

The efficacy of particular blocking reagents in preventing organ transplant rejection or GVHD can be assessed using animal models that are predictive of efficacy in humans. Examples of appropriate systems which can be used include allogeneic cardiac grafts in rats and xenogeneic pancreatic islet cell grafts in mice, both of which have been used to examine the immunosuppressive effects of CTLA4Ig fusion proteins in vivo as described in Lenschow et al., Science 257:789-792 (1992) and Turka et al., Proc. Natl. Acad. Sci USA, 89:11102-11105 (1992). In addition, murine models of GVHD (see Paul ed.,

Fundamental Immunology, Raven Press, New York, 1989, pp. 846-847) can be used to determine the effect of blocking B lymphocyte antigen function in vivo on the development of that disease.

Blocking antigen function may also be therapeutically useful for treating autoimmune diseases. Many autoimmune disorders are the result of inappropriate activation of T cells that are reactive against 5 self tissue and which promote the production of cytokines and autoantibodies involved in the pathology of the diseases. Preventing the activation of autoreactive T cells may reduce or eliminate disease symptoms. Administration of reagents which block costimulation of T cells by disrupting receptor/ligand interactions of B lymphocyte antigens can be used to inhibit T cell activation and prevent production of autoantibodies or T cell-derived cytokines which potentially involved in the disease 10 process. Additionally, blocking reagents may induce antigen-specific tolerance of autoreactive T cells which could lead to long-term relief from the disease. The efficacy of blocking reagents in preventing or alleviating autoimmune disorders can be determined using a number of well-characterized animal models of human autoimmune diseases. Examples include murine experimental autoimmune encephalitis, systemic lupus erythmatosis in MRL/pr/pr mice or NZB hybrid mice, murine autoimmuno 15 collagen arthritis, diabetes mellitus in OD mice and BB rats, and murine experimental myasthenia gravis (see Paul ed., Fundamental Immunology, Raven Press, New York, 1989, pp. 840-856).

Upregulation of an antigen function (preferably a B lymphocyte antigen function), as a means of up regulating immune responses, may also be useful in therapy. Upregulation of immune responses may involve either enhancing an existing immune response or eliciting an initial immune response as shown 20 by the following examples. For instance, enhancing an immune response through stimulating B lymphocyte antigen function may be useful in cases of viral infection. In addition, systemic viral diseases such as influenza, the common cold, and encephalitis might be alleviated by the administration of stimulatory form of B lymphocyte antigens systemically.

Alternatively, antiviral immune responses may be enhanced in an infected patient by removing 25 T cells from the patient, costimulating the T cells in vitro with viral antigen-pulsed APCs either expressing the proteins or polypeptides described above or together with a stimulatory form of the protein or polypeptide and reintroducing the *in vitro* primed T cells into the patient. The infected cells would now be capable of delivering a costimulatory signal to T cells in vivo, thereby activating the T cells.

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In another application, upregulation or enhancement of antigen function (preferably B lymphocyte antigen function) may be useful in the induction of tumor immunity. Tumor cells (e.g., sarcoma, melanoma, lymphoma, leukemia, neuroblastoma, carcinoma) transfected with one of the above-described nucleic acids encoding a gratein or polypeptide can be administered to a subject to evercome tumor-specific tolerance in the subject. If desired, the tumor cell can be transfected to express. 35 a combination of peptides. For example, tumor cells obtained from a patient can be transfected ex vivo with an expression vector directing the expression of a peptide having B7-2-like activity alone, or in conjunction with a peptide having B7-1-like activity and/or B7-3-like activity. The transfected tumor

cells are returned to the patient to result in expression of the peptides on the surface of the transfected cell. Alternatively, gene therapy techniques can be used to target a tumor cell for transfection in vivo.

The presence of the protein or polypeptide encoded by the nucleic acids described above having the activity of a B lymphocyte antigen(s) on the surface of the turnor cell provides the necessary 5 costimulation signal to T cells to induce a T cell mediated immune response against the transfected tumor cells. In addition, tumor cells which lack or which fail to reexpress sufficient amounts of MHC class I or MHC class II molecules can be transfected with nucleic acids encoding all or a portion of (e.g., a cytoplasmic-domain truncated portion) of an MHC class I α chain and β₂ microglobulin or an MHC class II α chain and an MHC class II β chain to thereby express MHC class I or MHC class II proteins 10 on the cell surface, respectively. Expression of the appropriate MHC class I or class II molecules in conjunction with a peptide having the activity of a B lymphocyte antigen (e.g., B7-1, B7-2, B7-3) induces a T cell mediated immune response against the transfected tumor cell. Optionally, a nucleic acid encoding an antisense construct which blocks expression of an MHC class II associated protein, such as the invariant chain, can also be cotransfected with a DNA encoding a protein or polypeptide having the 15 activity of a B lymphocyte antigen to promote presentation of tumor associated antigens and induce tumor specific immunity. Thus, the induction of a T cell mediated immune response in a human subject may be sufficient to overcome tumor-specific tolerance in the subject. Alternatively, as described in more detail below, nucleic acids encoding these immune system regulator proteins or polypeptides or nucleic acids regulating the expression of such proteins or polypeptides may be introduced into 20 appropriate host cells to increase or decrease the expression of the proteins as desired.

EXAMPLE 26

Assaying the Expressed Proteins or Polypeptides for Hematopoiesis Regulating Activity

25 The proteins or polypeptides encoded by the nucleic acids described above may also be evaluated for their hematopoiesis regulating activity. For example, the effect of the proteins or polypeptides on embryonic stem cell differentiation may be evaluated. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references: Johansson et al. Cell. Biol. 15:141-151, 1995; Keller et al., Mol. Cell. Biol. 13:473-486, 1993;
30 McClanahan et al., Blood 81:2903-2915, 1993.

The proteins or polypeptides encoded by the nucleic acids described above may also be evaluated for their influence on the lifetime of stem cells and stem cell differentiation. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references: Freshney, M.G. Methylcellulose Colony Forming Assays, in <u>Culture of Hematopoietic Cells</u>.

35 R.I. Freshney, et al. Eds. pp. 265-268, Wiley-Liss, Inc., New York, NY. 1994; Hirayama et al., Proc. Natl. Acad. Sci. USA 89:5907-5911, 1992; McNiece, I.K. and Briddell, R.A. Primitive Hematopoietic Colony Forming Cells with High Proliferative Potential, in <u>Culture of Hematopoietic Cells</u>. supra;

Neben et al., Experimental Hematology 22:353-359, 1994; Ploemacher, R.E. Cobblestone Area Forming Cell Assay, In Culture of Hematopoietic Cells, supra; Spooncer, E., Dexter, M. and Allen, T. Long Term Bone Marrow Cultures in the Presence of Stromal Cells, in Culture of Hematopoietic Cells supra; and Sutherland, H.J. Long Term Culture Initiating Cell Assay, in Culture of Hematopoietic Cells. supra.

Those proteins or polypeptides which exhibit hematopoiesis regulatory activity may then be formulated as pharmaceuticals and used to treat clinical conditions in which regulation of hematopoeisis is beneficial. For example, a protein or polypeptide of the present invention may be useful in regulation of hematopoiesis and, consequently, in the treatment of myeloid or lymphoid cell deficiencies. Even marginal biological activity in support of colony forming cells or of factor-dependent cell lines indicates 10 involvement in regulating hematopoiesis, e.g. in supporting the growth and proliferation of erythroid progenitor cells alone or in combination with other cytokines, thereby indicating utility, for example, in treating various anemias or for use in conjunction with irradiation/chemotherapy to stimulate the production of erythroid precursors and/or erythroid cells; in supporting the growth and proliferation of myeloid cells such as granulocytes and monocytes/macrophages (i.e., traditional CSF activity) useful, for 15 example, in conjunction with chemotherapy to prevent or treat consequent myelo-suppression; in supporting the growth and proliferation of megakaryocytes and consequently of platelets thereby allowing prevention or treatment of various platelet disorders such as thrombocytopenia, and generally for use in place of or complimentary to platelet transfusions; and/or in supporting the growth and proliferation of hematopoietic stem cells which are capable of maturing to any and all of the above-20 mentioned hematopoietic cells and therefore find therapeutic utility in various stem cell disorders (such as those usually treated with transplantion, including, without limitation, aplastic anemia and paroxysmal nocturnal hemoglobinuria), as well as in repopulating the stem cell compartment post irradiation/chemotherapy, either in-vivo or ex-vivo (i.e., in conjunction with bone marrow transplantation or with peripheral progenitor cell transplantation (homologous or heterologous)) as 25 normal cells or genetically manipulated for gene therapy. Alternatively, as described in more detail below, nucleic acids encoding these proteins or polypeptides or nucleic acids regulating the expression of these proteins or polypeptides may be introduced into appropriate host cells to increase or decrease the expression of the proteins as desired.

EXAMPLE 27 30

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Assaying the Expressed Proteins or Polypeptides

for Regulation of Tissue Growth

The proteins or polypeptides encoded by the nucleic acids described above may also be evaluated for their effect on tissue growth. Numerous assays for such activity are familiar to those _____ 35 skilled in the art, including the assays disclosed in International Patent Publication No. WO95/16035, International Patent Publication No. WO95/05846 and International Patent Publication No. WO91/07491.

Assays for wound healing activity include, without limitation, those described in: Winter, *Epidermal Wound Healing*, pps. 71-112 (Maibach, H1 and Rovee, DT, eds.), Year Book Medical Publishers, Inc., Chicago, as modified by Eaglstein and Mertz, J. Invest. Dermatol 71:382-84 (1978).

Those proteins or polypeptides which are involved in the regulation of tissue growth may then 5 be formulated as pharmaceuticals and used to treat clinical conditions in which regulation of tissue growth is beneficial. For example, a protein or polypeptide may have utility in compositions used for bone, cartilage, tendon, ligament and/or nerve tissue growth or regeneration, as well as for wound healing and tissue repair and replacement, and in the treatment of burns, incisions and ulcers.

A protein or polypeptide encoded by the nucleic acids described above which induces cartilage and/or bone growth in circumstances where bone is not normally formed, has application in the healing of bone fractures and cartilage damage or defects in humans and other animals. Such a preparation employing a protein or polypeptide of the invention may have prophylactic use in closed as well as open fracture reduction and also in the improved fixation of artificial joints. *De novo* bone synthesis induced by an osteogenic agent contributes to the repair of congenital, trauma induced, or oncologic resection induced craniofacial defects, and also is useful in cosmetic plastic surgery.

A protein or polypeptide of this invention may also be used in the treatment of periodontal disease, and in other tooth repair processes. Such agents may provide an environment to attract bone-forming cells, stimulate growth of bone-forming cells or induce differentiation of progenitors of bone-forming cells. A protein of the invention may also be useful in the treatment of osteoporosis or osteoarthritis, such as through stimulation of bone and/or cartilage repair or by blocking inflammation or processes of tissue destruction (collagenase activity, osteoclast activity, etc.) mediated by inflammatory processes.

Another category of tissue regeneration activity that may be attributable to the proteins or polypeptides encoded by the nucleic acids described above is tendon/ligament formation. A protein or polypeptide encoded by the nucleic acids described above, which induces tendon/ligament-like tissue or other tissue formation in circumstances where such tissue is not normally formed, has application in the healing of tendon or ligament tears, deformities and other tendon or ligament defects in humans and other animals. Such a preparation employing a tendon/ligament-like tissue inducing protein may have prophylactic use in preventing damage to tendon or ligament tissue, as well as use in the improved fixation of tendon or ligament to bone or other tissues, and in repairing defects to tendon or ligament tissue. De novo tendon/ligament-like tissue formation induced by a protein or polypeptide of the present invention contributes to the repair of tendon or ligaments defects of congenital, traumatic or other origin and is also useful in cosmetic plastic surgery for attachment or repair of tendons or ligaments. The proteins or polypeptides of the present invention may provide an environment to attract tendon-or ligament-forming cells, stimulate growth of tendon- or ligament-forming cells, induce differentiation of progenitors of tendon- or ligament-forming cells, or induce growth of tendon/ligament cells or

progenitors ex vivo for return in vivo to effect tissue repair. The proteins or polypeptides of the

invention may also be useful in the treatment of tendinitis, carpal tunnel syndrome and other tendon or ligament defects. The therapeutic compositions may also include an appropriate matrix and/or sequestering agent as a carrier as is well known in the art.

The proteins or polypeptides of the present invention may also be useful for proliferation of

5 neural cells and for regeneration of nerve and brain tissue, *i.e.*, for the treatment of central and peripheral
nervous system diseases and neuropathies, as well as mechanical and traumatic disorders, which involve
degeneration, death or trauma to neural cells or nerve tissue. More specifically, a protein or polypeptide
may be used in the treatment of diseases of the peripheral nervous system, such as peripheral nerve
injuries, peripheral neuropathy and localized neuropathies, and central nervous system diseases, such as
10 Alzheimer's, Parkinson's disease, Huntington's disease, amyotrophic lateral sclerosis, and Shy-Drager
syndrome. Further conditions which may be treated in accordance with the present invention include
mechanical and traumatic disorders, such as spinal cord disorders, head trauma and cerebrovascular
diseases such as stroke. Peripheral neuropathies resulting from chemotherapy or other medical therapies
may also be treatable using a protein or polypeptide of the invention.

Proteins or polypeptides of the invention may also be useful to promote better or faster closure of non-healing wounds, including without limitation pressure ulcers, ulcers associated with vascular insufficiency, surgical and traumatic wounds, and the like.

It is expected that a protein or polypeptide of the present invention may also exhibit activity for generation or regeneration of other tissues, such as organs (including, for example, pancreas, liver, intestine, kidney, skin, endothelium) muscle (smooth, skeletal or cardiac) and vascular (including vascular endothelium) tissue, or for promoting the growth of cells comprising such tissues. Part of the desired effects may be by inhibition or modulation of fibrotic scarring to allow normal tissue to generate. A protein or polypeptide of the invention may also exhibit angiogenic activity.

A protein or polypeptide of the present invention may also be useful for gut protection or regeneration and treatment of lung or liver fibrosis, reperfusion injury in various tissues, and conditions resulting from systemic cytokine damage.

A protein or polypeptide of the present invention may also be useful for promoting or inhibiting differentiation of tissues described above from precursor tissues or cells; or for inhibiting the growth of tissues described above.

Alternatively, as described in more detail below, nucleic acids encoding tissue growth regulating activity proteins or polypeptides or nucleic acids regulating the expression of such proteins or polypeptides may be introduced into appropriate host cells to increase or decrease the expression of the proteins as desired.

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The proteins or polypeptides of the present invention may also be evaluated for their ability to regulate reproductive hormones, such as follicle stimulating hormone. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references: Vale et al., Endocrinol. 91:562-572, 1972; Ling et al., Nature 321:779-782, 1986; Vale et al., Nature 5 321:776-779, 1986; Mason et al., Nature 318:659-663, 1985; Forage et al., Proc. Natl. Acad. Sci. USA 83:3091-3095, 1986. Chapter 6.12 in Current Protocols in Immunology, J.E. Coligan et al. Eds. Greene Publishing Associates and Wiley-Intersciece; Taub et al. J. Clin. Invest. 95:1370-1376, 1995; Lind et al. APMIS 103:140-146, 1995; Muller et al. Eur. J. Immunol: 25:1744-1748; Gruber et al. J. Immunol. 152:5860-5867, 1994; Johnston et al., J Immunol. 153:1762-1768, 1994.

Those proteins or polypeptides which exhibit activity as reproductive hormones or regulators of cell movement may then be formulated as pharmaceuticals and used to treat clinical conditions in which regulation of reproductive hormones are beneficial. For example, a protein or polypeptide may exhibit activin- or inhibin-related activities. Inhibins are characterized by their ability to inhibit the release of follicle stimulating hormone (FSH), while activins are characterized by their ability to stimulate the 15 release of FSH. Thus, a protein or polypeptide of the present invention, alone or in heterodimers with a member of the inhibin family, may be useful as a contraceptive based on the ability of inhibins to decrease fertility in female mammals and decrease spermatogenesis in male mammals. Administration of sufficient amounts of other inhibins can induce infertility in these mammals. Alternatively, the protein or polypeptide of the invention, as a homodimer or as a heterodimer with other protein subunits 20 of the inhibin-B group, may be useful as a fertility inducing therapeutic, based upon the ability of activin molecules in stimulating FSH release from cells of the anterior pituitary. See, for example, United States Patent 4,798,885. A protein or polypeptide of the invention may also be useful for advancement of the onset of fertility in sexually immature mammals, so as to increase the lifetime reproductive performance of domestic animals such as cows, sheep and pigs.

Alternatively, as described in more detail below, nucleic acids encoding reproductive hormone regulating activity proteins or polypeptides or nucleic acids regulating the expression of such proteins or polypeptides may be introduced into appropriate host cells to increase or decrease the expression of the proteins or polypeptides as desired.

EXAMPLE 29 30

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Assaying the Expressed Proteins or Polypeptides For Chemotactic/Chemokinetic Activity The proteins or polypeptides of the present invention may also be evaluated for chemotactic/chemokinetic activity. For example, a protein or polypeptide of the present invention may have chemotactic or chemokinetic activity (e.g., act as a chemokine) for mammalian cells, including, for 35 example, monocytes, fibroblasts, neutrophils, T-cells, mast cells, eosinophils, epithelial and/or endothelial cells. Chemotactic and chemokinetic proteins or polypeptides can be used to mobilize or attract a desired cell population to a desired site of action. Chemotactic or chemokinetic proteins or

polypeptides provide particular advantages in treatment of wounds and other trauma to tissues, as well as in treatment of localized infections. For example, attraction of lymphocytes, monocytes or neutrophils to tumors or sites of infection may result in improved immune responses against the tumor or infecting agent.

A protein or polypeptide has chemotactic activity for a particular cell population if it can stimulate, directly or indirectly, the directed orientation or movement of such cell population.

Preferably, the protein or polypeptide has the ability to directly stimulate directed movement of cells.

Whether a particular protein or polypeptide has chemotactic activity for a population of cells can be readily determined by employing such protein or polypeptide in any known assay for cell chemotaxis.

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The activity of a protein or polypeptide of the invention may, among other means, be measured by the following methods:

Assays for chemotactic activity (which will identify proteins or polypeptides that induce or prevent chemotaxis) consist of assays that measure the ability of a protein or polypeptide to induce the migration of cells across a membrane as well as the ability of a protein or polypeptide to induce the adhesion of one cell population to another cell population. Suitable assays for movement and adhesion include, without limitation, those described in: *Current Protocols in Immunology*, Ed by J.E. Coligan, A.M. Kruisbeek, D.H. Margulies, E.M. Shevach, W. Strober, Pub. Greene Publishing Associates and Wiley-Interscience, Chapter 6.12: 6.12.1-6.12.28; Taub *et al. J. Clin. Invest.* 95:1370-1376, 1995; Lind *et al. APMIS* 103:140-146, 1995; Mueller *et al., Eur. J. Immunol.* 25:1744-1748; Gruber *et al. J. Immunol.* 152:5860-5867, 1994; Johnston *et al. J. Immunol.*, 153:1762-1768, 1994.

EXAMPLE 30

Assaying the Expressed Proteins or Polypeptides for Regulation of Blood Clotting

The proteins or polypeptides of the present invention may also be evaluated for their effects on blood clotting. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references: Linet et al., J. Clin. Pharmacol. 26:131-140, 1986; Burdick et al., Thrombosis Res. 45:413-419, 1987; Humphrey et al., Fibrinolysis 5:71-79 (1991); Schaub, Prostaglandins 35:467-474, 1988.

Those proteins or polypeptides which are involved in the regulation of blood clotting may then

be formulated as pharmaceuticals and used to treat clinical conditions in which regulation of blood

clotting is beneficial. For example, a protein or polypeptide of the invention may also exhibit hemostatic

or thrombolytic activity. As a result, such a protein or polypeptide is expected to be useful in treatment

of various coagulations disorders (including hereditary disorders, such as hemophilias) or to enhance

coagulation and other hemostatic events in treating wounds resulting from trauma, surgery or other

causes. A protein or polypeptide of the invention may also be useful for dissolving or inhibiting

formation of thromboses and for treatment and prevention of conditions resulting therefrom (such as

infarction of cardiac and central nervous system vessels (e.g., stroke)). Alternatively, as described in

more detail below, nucleic acids encoding blood clotting activity proteins or polypeptides or nucleic acids regulating the expression of such proteins or polypeptides may be introduced into appropriate host cells to increase or decrease the expression of the proteins or polypeptides as desired.

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EXAMPLE 31

Assaying the Expressed Proteins or Polypeptides for Involvement in Receptor/Ligand Interactions

The proteins or polypeptides of the present invention may also be evaluated for their involvement in receptor/ligand interactions. Numerous assays for such involvement are familiar to those 10 skilled in the art, including the assays disclosed in the following references: Chapter 7. 7.28.1-7.28.22) in Current Protocols in Immunology, J.E. Coligan et al. Eds. Greene Publishing Associates and Wiley-Interscience; Takai et al., Proc. Natl. Acad. Sci. USA 84:6864-6868, 1987; Bierer et al., J. Exp. Med. 168:1145-1156, 1988; Rosenstein et al., J. Exp. Med. 169:149-160, 1989; Stoltenborg et al., J. Immunol. Methods 175:59-68, 1994; Stitt et al., Cell 80:661-670, 1995; Gyuris et al., Cell 75:791-803, 1993.

For example, the proteins or polypeptides of the present invention may also demonstrate activity as receptors, receptor ligands or inhibitors or agonists of receptor/ligand interactions. Examples of such receptors and ligands include, without limitation, cytokine receptors and their ligands, receptor kinases and their ligands, receptor phosphatases and their ligands, receptors involved in cell-cell interactions and their ligands (including without limitation, cellular adhesion molecules (such as selectins, integrins and 20 their ligands) and receptor/ligand pairs involved in antigen presentation, antigen recognition and development of cellular and humoral immune responses). Receptors and ligands are also useful for screening of potential peptide or small molecule inhibitors of the relevant receptor/ligand interaction. A protein or polypeptide of the present invention (including, without limitation, fragments of receptors and ligands) may be useful as inhibitors of receptor/ligand interactions. Alternatively, as described in more 25 detail below, nucleic acids encoding proteins or polypeptides involved in receptor/ligand interactions or nucleic acids regulating the expression of such proteins or polypeptides may be introduced into appropriate host cells to increase or decrease the expression of the proteins or polypeptides as desired.

EXAMPLE 32

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Assaying the Proteins or Polypeptides for Anti-Inflammatory Activity

The proteins or polypeptides of the present invention may also be evaluated for antiinflammatory activity. The anti-inflammatory activity may be achieved by providing a stimulus to cells involved in the inflammatory response, by inhibiting or promoting cell-cell interactions (such as, for example cell adhesion), by inhibiting or promoting chemotaxis of cells involved in the 35 inflammatory process, inhibiting or promoting cell extravasation, or by stimulating or suppressing production of other factors which more directly inhibit or promote an inflammatory response. Proteins or polypeptides exhibiting such activities can be used to treat inflammatory conditions

including chronic or acute conditions, including without limitation inflammation associated with infection (such as septic shock, sepsis or systemic inflammatory response syndrome), ischemia-reperfusioninury, endotoxin lethality, arthritis, complement-mediated hyperacute rejection, nephritis, cytokine- or chemokine-induced lung injury, inflammatory bowel disease, Crohn's disease or resulting from over production of cytokines such as TNF or IL-1. Proteins or polypeptides of the invention may also be useful to treat anaphylaxis and hypersensitivity to an antigenic substance or material. Alternatively, as described in more detail below, nucleic acids encoding anti-inflammatory activity proteins or polypeptides or nucleic acids regulating the expression of such proteins or polypeptides may be introduced into appropriate host cells to increase or decrease the expression of the proteins or polypeptides as desired.

EXAMPLE 33

Assaying the Expressed Proteins or Polypeptides for Tumor Inhibition Activity

The proteins or polypeptides of the present invention may also be evaluated for tumor inhibition activity. In addition to the activities described above for immunological treatment or prevention of tumors, a protein or polypeptide of the invention may exhibit other anti-tumor activities. A protein or polypeptide may inhibit tumor growth directly or indirectly (such as, for example, via ADCC). A protein or polypeptide may exhibit its tumor inhibitory activity by acting on tumor tissue or tumor precursor tissue, by inhibiting formation of tissues necessary to support tumor growth (such as, for example, by inhibiting angiogenesis), by causing production of other factors, agents or cell types which inhibit tumor growth, or by suppressing, eliminating or inhibiting factors, agents or cell types which promote tumor growth. Alternatively, as described in more detail below, nucleic acids encoding proteins or polypeptides with tumor inhibition activity or nucleic acids regulating the expression of such proteins or polypeptides may be introduced into appropriate host cells to increase or decrease the expression of the proteins or polypeptides as desired.

A protein or polypeptide of the invention may also exhibit one or more of the following additional activities or effects: inhibiting the growth, infection or function of, or killing, infectious agents, including, without limitation, bacteria, viruses, fungi and other parasites; effecting (suppressing or enhancing) bodily characteristics, including, without limitation, height, weight, hair color, eye color, skin, fat to lean ratio or other tissue pigmentation, or organ or body part size or shape (such as, for example, breast augmentation or diminution, change in bone form or shape); effecting biorhythms or circadian cycles or rhythms; effecting the fertility of male or female subjects; effecting the metabolism, catabolism, anabolism, processing, utilization, storage or dimination of dietary fat, lipid, protein, carbohydrate, vitamins, minerals, cofactors or other nutritional factors or component(s); effecting behavioral characteristics, including, without limitation, appetite, libido, stress, cognition (including cognitive disorders), depression (including depressive disorders) and violent behaviors; providing analgesic effects or other pain reducing effects; promoting differentiation and growth of embryonic stem

cells in lineages other than hematopoietic lineages; hormonal or endocrine activity; in the case of enzymes, correcting deficiencies of the enzyme and treating deficiency-related diseases; treatment of hyperproliferative disorders (such as, for example, psoriasis); immunoglobulin-like activity (such as, for example, the ability to bind antigens or complement); and the ability to act as an antigen in a vaccine composition to raise an immune response against such protein or another material or entity which is cross-reactive with such protein. Alternatively, as described in more detail below, nucleic acids encoding proteins or polypeptides involved in any of the above mentioned activities or nucleic acids regulating the expression of such proteins may be introduced into appropriate host cells to increase or decrease the expression of the proteins or polypeptides as desired.

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EXAMPLE 34

Identification of Proteins or Polypeptides which Interact with Proteins or Polypeptides of the Present Invention

15 invention, such as receptor proteins, may be identified using two hybrid systems such as the Matchmaker Two Hybrid System 2 (Catalog No. K1604-1, Clontech). As described in the manual accompanying the kit, nucleic acids encoding the proteins or polypeptides of the present invention, are inserted into an expression vector such that they are in frame with DNA encoding the DNA binding domain of the yeast transcriptional activator GAL4. cDNAs in a cDNA library which encode proteins or polypeptides which might interact with the proteins or polypeptides of the present invention are inserted into a second expression vector such that they are in frame with DNA encoding the activation domain of GAL4. The two expression plasmids are transformed into yeast and the yeast are plated on selection medium which selects for expression of selectable markers on each of the expression vectors as well as GAL4 dependent expression of the HIS3 gene. Transformants capable of growing on medium lacking histidine are screened for GAL4 dependent lacZ expression. Those cells which are positive in both the histidine selection and the lacZ assay contain plasmids encoding proteins or polypeptides which interact with the proteins or polypeptides of the present invention.

Alternatively, the system described in Lustig et al., Methods in Enzymology 283: 83-99 (1997) may be used for identifying molecules which interact with the proteins or polypeptides of the present invention. In such systems, in vitro transcription reactions are performed on a pool of vectors containing nucleic acid inserts which encode the proteins or polypeptides of the present invention. The nucleic acid inserts are cloned downstream of a promoter which drives in vitro transcription. The resulting pools of mRNAs are introduced into Xenopus laevis oocytes. The oocytes are then assayed for a desired activity.

Alternatively, the pooled in vitro transcription products produced as described above may be translated in vitro. The pooled in vitro translation products can be assayed for a desired activity or for interaction with a known protein or polypeptide.

Proteins, polypeptides or other molecules interacting with proteins or polypeptides of the present invention can be found by a variety of additional techniques. In one method, affinity columns containing the protein or polypeptide of the present invention can be constructed. In some versions, of this method the affinity column contains chimeric proteins in which the protein or polypeptide of the present invention is fused to glutathione S-transferase. A mixture of cellular proteins or pool of expressed proteins as described above and is applied to the affinity column. Molecules interacting with the protein or polypeptide attached to the column can then be isolated and analyzed on 2-D electrophoresis gel as described in Ramunsen et al. Electrophoresis, 18, 588-598 (1997). Alternatively, the molecules retained on the affinity column can be purified by electrophoresis based methods and sequenced. The same method can be used to isolate antibodies, to screen phage display products, or to screen phage display human antibodies.

Molecules interacting with the proteins or polypeptides of the present invention can also be screened by using an Optical Biosensor as described in Edwards & Leatherbarrow, Analytical Biochemistry, 246, 1-6 (1997). The main advantage of the method is that it allows the determination of the association rate between the protein or polypeptide and other interacting molecules. Thus, it is possible to specifically select interacting molecules with a high or low association rate. Typically a target molecule is linked to the sensor surface (through a carboxymethl dextran matrix) and a sample of test molecules is placed in contact with the target molecules. The binding of a test molecule to the target molecule causes a change in the refractive index and/ or thickness. This change is detected by the Biosensor provided it occurs in the evanescent field (which extends a few hundred nanometers from the sensor surface). In these screening assays, the target molecule can be one of the proteins or polypeptides of the present invention and the test sample can be a collection of proteins, polypeptides or other molecules extracted from tissues or cells, a pool of expressed proteins, combinatorial peptide and/ or chemical libraries, or phage displayed peptides. The tissues or cells from which the test

In other methods, a target protein or polypeptide is immobilized and the test population is a collection of unique proteins or polypeptides of the present invention.

To study the interaction of the proteins or polypeptides of the present invention with drugs, the microdialysis coupled to HPLC method described by Wang et al., Chromatographia, 44, 205-208(1997) or the affinity capillary electrophoresis method described by Busch et al., J. Chromatogr. 777:311-328 (1997)can be used.

The system described in U.S. Patent No. 5,654,150 may also be used to identify molecules which interact with the proteins or polypeptides of the present invention. In this system, pools of nucleis acids encoding the proteins or polypeptides of the present invention are transcribed and translated in vitro and the reaction products are assayed for interaction with a known polypeptide or antibody.

It will be appreciated by those skilled in the art that the proteins or polypeptides of the present invention may be assayed for numerous activities in addition to those specifically enumerated above.

For example, the expressed proteins or polypeptides may be evaluated for applications involving control and regulation of inflammation, tumor proliferation or metastasis, infection, or other clinical conditions. In addition, the proteins or polypeptides may be useful as nutritional agents or cosmetic agents.

The proteins or polypeptides of the present invention may be used to generate antibodies

5 capable of specifically binding to the proteins or polypeptides of the present invention. The
antibodies may be monoclonal antibodies or polyclonal antibodies. As used herein, "antibody" refers
to a polypeptide or group of polypeptides which are comprised of at least one binding domain, where
a binding domain is formed from the folding of variable domains of an antibody molecule to form
three-dimensional binding spaces with an internal surface shape and charge distribution

10 complementary to the features of an antigenic determinant of an antigen., which allows an
immunological reaction with the antigen. Antibodies include recombinant proteins comprising the
binding domains, as wells as fragments, including Fab, Fab', F(ab)2, and F(ab')2 fragments.

As used herein, an "antigenic determinant" is the portion of an antigen molecule, that determines the specificity of the antigen-antibody reaction. An "epitope" refers to an antigenic determinant of a polypeptide. An epitope can comprise as few as 3 amino acids in a spatial conformation which is unique to the epitope. Generally an epitope consists of at least 6 such amino acids, and more usually at least 8-10 such amino acids. Methods for determining the amino acids which make up an epitope include x-ray crystallography, 2-dimensional nuclear magnetic resonance, and epitope mapping e.g. the Pepscan method described by H. Mario Geysen et al. 1984. Proc. Natl. Acad. Sci. U.S.A. 81:3998-4002; PCT Publication No. WO 84/03564; and PCT Publication No. WO 84/03506.

In some embodiments, the antibodies may be capable of specifically binding to a protein or polypeptide encoded by EST-related nucleic acids, fragments of EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids.

In some embodiments, the antibody may be capable of binding an antigenic determinant or an epitope in a protein or polypeptide encoded by EST-related nucleic acids, fragments of EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids.

In other embodiments, the antibodies may be capable of specifically binding to an EST-related polypeptide, fragment of an EST-related polypeptide, positional segment of an EST-related polypeptide or fragment of a positional segment of an EST-related polypeptide. In some embodiments, the antibody may be capable of binding an antigenic determinant or an epitope in an EST-related polypeptide, tragment of an EST-related polypeptide, positional segment of an EST-related polypeptide or fragment of a positional segment of an EST-related polypeptide.

In the case of secreted proteins, the antibodies may be capable of binding a full-length protein encoded by a nucleic acid of the present invention, a mature protein (i.e. the protein generated by

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cleavage of the signal peptide) encoded by a nucleic acid of the present invention, or a signal peptide encoded by a nucleic acid of the present invention.

EXAMPLE 35

Production of an Antibody to a Human Polypeptide or Protein

The above described EST-related nucleic acids, fragments of EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids or nucleic acids encoding EST-related polypeptides, fragments of EST-related polypeptides, positional segments of EST-related polypeptides or fragments of positional segments of 10 EST-related polypeptides are operably linked to promoters and introduced into cells as described above.

In the case of secreted proteins, nucleic acids encoding the full protein (i.e. the mature protein and the signal peptide), nucleic acids encoding the mature protein (i.e. the protein generated by cleavage of the signal peptide), or nucleic acids encoding the signal peptide are operably linked to promoters and introduced into cells as described above.

The encoded proteins or polypeptides are then substantially purified or isolated as described above. The concentration of protein in the final preparation is adjusted, for example, by concentration on an Amicon filter device, to the level of a few µg/ml. Monoclonal or polyclonal antibody to the protein or polypeptide can then be prepared as follows:

1. Monoclonal Antibody Production by Hybridoma Fusion

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Monoclonal antibody to epitopes of any of the proteins or polypeptides identified and isolated as described can be prepared from murine hybridomas according to the classical method of Kohler, and Milstein, Nature 256:495 (1975) or derivative methods thereof. Briefly, a mouse is repetitively inoculated with a few micrograms of the selected protein or peptides derived therefrom over a period of a few weeks. The mouse is then sacrificed, and the antibody producing cells of the spleen isolated. The 25 spleen cells are fused by means of polyethylene glycol with mouse myeloma cells, and the excess unfused cells destroyed by growth of the system on selective media comprising aminopterin (HAT media). The successfully fused cells are diluted and aliquots of the dilution placed in wells of a microtiter plate where growth of the culture is continued. Antibody-producing clones are identified by detection of antibody in the supernatant fluid of the wells by immunoassay procedures, such as Elisa, as 30 originally described by Engvall, Meth. Enzymol. 70:419 (1980). Selected positive clones can be expanded and their monoclonal antibody product harvested for use. Detailed procedures for monoclonal antibody production are described in Davis, L. et al. in Basic Methods in Molecular Biology Elsevier, New York. Section 21-2.

2. Polyclonal Antibody Production by Immunization

Polyclonal antiserum containing antibodies to heterogenous epitopes of a single protein or polypeptide can be prepared by immunizing suitable animals with the expressed protein or peptides derived therefrom, which can be unmodified or modified to enhance immunogenicity. Effective

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polyclonal antibody production is affected by many factors related both to the antigen and the host species. For example, small molecules tend to be less immunogenic than others and may require the use of carriers and adjuvant. Also, host animals response vary depending on site of inoculations and doses, with both inadequate or excessive doses of antigen resulting in low titer antisera. Small doses (ng level) of antigen administered at multiple intradermal sites appears to be most reliable. An effective immunization protocol for rabbits can be found in Vaitukaitis. et al.J. Clin. Endocrinol. Metab. 33:988-991 (1971).

Booster injections can be given at regular intervals, and antiserum harvested when antibody titer thereof, as determined semi-quantitatively, for example, by double immunodiffusion in agar against look nown concentrations of the antigen, begins to fall. See, for example, Ouchterlony, et al., Chap. 19 in: Handbook of Experimental Immunology D. Wier (ed) Blackwell (1973). Plateau concentration of antibody is usually in the range of 0.1 to 0.2 mg/ml of serum (about 12 μM). Affinity of the antisera for the antigen is determined by preparing competitive binding curves, as described, for example, by Fisher, D., Chap. 42 in: Manual of Clinical Immunology, 2d Ed. (Rose and Friedman, Eds.) Amer. Soc. For Microbiol., Washington, D.C. (1980).

Antibody preparations prepared according to either of the above protocols are useful in a variety of contexts. In particular, the antibodies may be used in immunoaffinity chromatography techniques such as those described below to facilitate large scale isolation, purification, or enrichment of the proteins or polypeptides encoded by EST-related nucleic acids, positional segments of EST-related nucleic acids or for the isolation, purification or enrichment of EST-related polypeptides, fragments of EST-related polypeptides, positional segments of EST-related polypeptides.

In the case of secreted proteins, the antibodies may be used for the isolation, purification, or enrichment of the full protein (*i.e.* the mature protein and the signal peptide), the mature protein (*i.e.* the protein generated by cleavage of the signal peptide), or the signal peptide are operably linked to promoters and introduced into cells as described above.

Additionally, the antibodies may be used in immunoaffinity chromatography techniques such as those described below to isolate, purify, or enrich polypeptides which have been linked to the proteins or polypeptides encoded by EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids or to isolate, purify, or enrich EST-related polypeptides, fragments of EST-related polypeptides, positional segments of EST-related polypeptides or fragments of positional segments of EST-related polypeptides.

The antibodies may also be used to determine the cellular localization of polypeptides encoded by the proteins or polypeptides encoded by EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids or the cellular

localization of EST-related polypeptides, fragments of EST-related polypeptides, positional segments of EST-related polypeptides or fragments of positional segments of EST-related polypeptides.

In addition, the antibodies may also be used to determine the cellular localization of polypeptides which have been linked to the proteins or polypeptides encoded by EST-related nucleic 5 acids, positional segments of EST-related nucleic acids or fragments of positional segments of ESTrelated nucleic acids or polypeptides which have been linked to EST-related polypeptides, fragments of EST-related polypeptides, positional segments of EST-related polypeptides or fragments of positional segments of EST-related polypeptides.

The antibodies may also be used in quantitative immunoassays which determine concentrations 10 of antigen-bearing substances in biological samples; they may also used semi-quantitatively or qualitatively to identify the presence of antigen in a biological sample or to identify the type of tissue present in a biological sample. The antibodies may also be used in therapeutic compositions for killing cells expressing the protein or reducing the levels of the protein in the body.

15 VI. Use of 5'ESTs or Consensus Contigated 5' ESTs or Sequences Obtainable Therefrom or **Portions Thereof as Reagents**

The EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids may be used as reagents in isolation procedures, diagnostic assays, and forensic procedures. For example, sequences from the EST-related nucleic acids, 20 positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids, may be detectably labeled and used as probes to isolate other sequences capable of hybridizing to them. In addition, the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids may be used to design PCR primers to be used in isolation, diagnostic, or forensic procedures.

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1. Use of EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids in isolation, diagnostic and forensic procedures

EXAMPLE 36

Preparation of PCR Primers and Amplification of DNA

The EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids may be used to prepare PCR primers for a variety of applications, including isolation procedures for cloning nucleic acids capable of hybridizing to such sequences, diagnostic techniques and forensic techniques. In some embodiments, the PCR primers at least 10, 15, 18, 20, 23, 25, 28, 30, 40, or 50 nucleotides in length. In some embodiments, the PCR 35 primers may be more than 30 bases in length. It is preferred that the primer pairs have approximately the same G/C ratio, so that melting temperatures are approximately the same. A variety of PCR techniques are familiar to those skilled in the art. For a review of PCR technology, see Molecular Cloning to

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Genetic Engineering White, B.A. Ed. in Methods in Molecular Biology 67: Humana Press, Totowa 1997. In each of these PCR procedures, PCR primers on either side of the nucleic acid sequences to be amplified are added to a suitably prepared nucleic acid sample along with dNTPs and a thermostable polymerase such as Taq polymerase, Pfu polymerase, or Vent polymerase. The nucleic acid in the sample is denatured and the PCR primers are specifically hybridized to complementary nucleic acid sequences in the sample. The hybridized primers are extended. Thereafter, another cycle of denaturation, hybridization, and extension is initiated. The cycles are repeated multiple times to produce an amplified fragment containing the nucleic acid sequence between the primer sites.

10 EXAMPLE 37

Use of the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids as probes

Probes derived from EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids may be labeled with detectable labels

15 familiar to those skilled in the art, including radioisotopes and non-radioactive labels, to provide a detectable probe. The detectable probe may be single stranded or double stranded and may be made using techniques known in the art, including *in vitro* transcription, nick translation, or kinase reactions. A nucleic acid sample containing a sequence capable of hybridizing to the labeled probe is contacted with the labeled probe. If the nucleic acid in the sample is double stranded, it may be denatured prior to contacting the probe. In some applications, the nucleic acid sample may be immobilized on a surface such as a nitrocellulose or nylon membrane. The nucleic acid sample may comprise nucleic acids obtained from a variety of sources, including genomic DNA, cDNA libraries, RNA, or tissue samples.

Procedures used to detect the presence of nucleic acids capable of hybridizing to the detectable probe include well known techniques such as Southern blotting, Northern blotting, dot blotting, colony hybridization, and plaque hybridization. In some applications, the nucleic acid capable of hybridizing to the labeled probe may be cloned into vectors such as expression vectors, sequencing vectors, or *in vitro* transcription vectors to facilitate the characterization and expression of the hybridizing nucleic acids in the sample. For example, such techniques may be used to isolate and clone sequences in a genomic library or cDNA library which are capable of hybridizing to the detectable probe as described in Example 20 above.

PCR primers made as described in Example 36 above may be used in forensic analyses, such as the DNA fingerprinting techniques described in Examples 38-42 below. Such analyses may utilize detectable probes or primers based on the sequences of the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids.

In one exemplary method, DNA samples are isolated from forensic specimens of, for example, hair, semen, blood or skin cells by conventional methods. A panel of PCR primers based on a number of the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids is then utilized in accordance with Example 36 to amplify DNA of approximately 100-200 bases in length from the forensic specimen. Corresponding sequences are obtained from a test subject. Each of these identification DNAs is then sequenced using standard techniques, and a simple database comparison determines the differences, if any, between the sequences from the subject and those from the sample. Statistically significant differences between the suspect's DNA sequences and those from the sample conclusively prove a lack of identity. This lack of identity can be proven, for example, with only one sequence. Identity, on the other hand, should be demonstrated with a large number of sequences, all matching. Preferably, a minimum of 50 statistically identical sequences of 100 bases in length are used to prove identity between the suspect and the sample.

EXAMPLE 39

Positive Identification by DNA Sequencing

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The technique outlined in the previous example may also be used on a larger scale to provide a unique fingerprint-type identification of any individual. In this technique, primers are prepared from a large number of EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids. Preferably, 20 to 50 different primers are used. These primers are used to obtain a corresponding number of PCR-generated DNA segments from the individual in question in accordance with Example 34. Each of these DNA segments is sequenced, using the methods set forth in Example 36. The database of sequences generated through this procedure uniquely identifies the individual from whom the sequences were obtained. The same panel of primers may then be used at any later time to absolutely correlate tissue or other biological specimen with that individual.

EXAMPLE 40

Southern Blot Forensic Identification

The procedure of Example 38 is repeated to obtain a panel of at least 10 amplified sequences

from an individual and a specimen. Preferably, the panel contains at least 50 amplified sequences. More preferably, the panel contains 100 amplified sequences. In some embodiments, the panel contains 200 amplified sequences. This PCR-generated DNA is then digested with one or a combination of, preferably, four base specific restriction enzymes. Such enzymes are commercially available and known to those of skill-in-the art. After-digestion, the resultant gene fragments are size separated in multiple.

duplicate wells on an agarose gel and transferred to nitrocellulose using Southern blotting techniques well known to those with skill in the art. For a review of Southern blotting see Davis et al. (Basic Methods in Molecular Biology, 1986, Elsevier Press. pp 62-65).

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A panel of probes based on the sequences of the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids are radioactively or colorimetrically labeled using methods known in the art, such as nick translation or end labeling, and hybridized to the Southern blot using techniques known in the art (Davis et al., supra).

5 Preferably, the probe is at least 10, 12, 15, 18, 20, 25, 28, 30, 35, 40, 50, 75, 100, 150, 200, 300, 400 or 500 nucleotides in length. Preferably, the probes are at least 10, 12, 15, 18, 20, 25, 28, 30, 35, 40, 50, 75, 100, 150, 200, 300, 400 or 500 nucleotides in length. In some embodiments, the probes are oligonucleotides which are 40 nucleotides in length or less.

Preferably, at least 5 to 10 of these labeled probes are used, and more preferably at least about 20 or 30 are used to provide a unique pattern. The resultant bands appearing from the hybridization of a large sample of EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids will be a unique identifier. Since the restriction enzyme cleavage will be different for every individual, the band pattern on the Southern blot will also be unique. Increasing the number of probes will provide a statistically higher level of confidence in the identification since there will be an increased number of sets of bands used for identification.

EXAMPLE 41

Dot Blot Identification Procedure

Another technique for identifying individuals using the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids disclosed herein utilizes a dot blot hybridization technique.

Genomic DNA is isolated from nuclei of subject to be identified. Probes are prepared that correspond to at least 10, preferably 50 sequences from the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids.

- 25 The probes are used to hybridize to the genomic DNA through conditions known to those in the art. The oligonucleotides are end labeled with P³² using polynucleotide kinase (Pharmacia). Dot Blots are created by spotting the genomic DNA onto nitrocellulose or the like using a vacuum dot blot manifold (BioRad, Richmond California). The nitrocellulose filter containing the genomic sequences is baked or UV linked to the filter, prehybridized and hybridized with labeled probe using techniques known in the art (Davis et al., supra). The ³²P labeled DNA fragments are sequentially hybridized with successively stringent
 - Tetramethylammonium chloride is useful for identifying clones containing small numbers of nucleotide mismatches (Wood et ai., Proc. Natl. Acad. Sci. USA 82(6):1385-3382 (1985)). A unique pattern of dots distinguishes one individual from another individual.

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EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids can be used as probes in the following alternative

conditions to detect minimal differences between the 30 bp sequence and the DNA.

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fingerprinting technique. In some embodiments, the probes are oligonucleotides which are 40 nucleotides in length or less.

Preferably, a plurality of probes having sequences from different EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related 5 nucleic acids are used in the alternative fingerprinting technique. Example 42 below provides a representative alternative fingerprinting procedure in which the probes are derived from EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids.

10 **EXAMPLE 42**

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Alternative "Fingerprint" Identification Technique

Oligonucleotides are prepared from a large number, e.g. 50, 100, or 200, EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of ESTrelated nucleic acids using commercially available oligonucleotide services such as Genset, Paris, 15 France. Preferably, the oligonucleotides are at least 10, 15, 18, 20, 23, 25 28, or 30 nucleotides in length. However, in some embodiments, the oligonucleotides may be more than 40, 50, 60 or 70 nucleotides in length.

Cell samples from the test subject are processed for DNA using techniques well known to those with skill in the art. The nucleic acid is digested with restriction enzymes such as EcoRI and XbaI. 20 Following digestion, samples are applied to wells for electrophoresis. The procedure, as known in the art, may be modified to accommodate polyacrylamide electrophoresis, however in this example, samples containing 5 ug of DNA are loaded into wells and separated on 0.8% agarose gels. The gels are transferred onto nitrocellulose using standard Southern blotting techniques.

10 ng of each of the oligonucleotides are pooled and end-labeled with P³². The nitrocellulose is 25 prehybridized with blocking solution and hybridized with the labeled probes. Following hybridization and washing, the nitrocellulose filter is exposed to X-Omat AR X-ray film. The resulting hybridization pattern will be unique for each individual.

It is additionally contemplated within this example that the number of probe sequences used can be varied for additional accuracy or clarity.

In addition to their applications in forensics and identification, EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids may be mapped to their chromosomal locations. Example 41 below describes radiation hybrid (RH) mapping of human chromosomal regions using EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids 35 Example 42 below describes a representative procedure for mapping EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids to their locations on human chromosomes. Example 43 below describes mapping of

EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids on metaphase chromosomes by Fluorescence In Situ Hybridization (FISH).

5 <u>2. Use of EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids in Chromosome Mapping</u>

EXAMPLE 43

Radiation hybrid mapping of EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of

10 <u>EST-related nucleic acids to the human genome</u>

Radiation hybrid (RH) mapping is a somatic cell genetic approach that can be used for high resolution mapping of the human genome. In this approach, cell lines containing one or more human chromosomes are lethally irradiated, breaking each chromosome into fragments whose size depends on the radiation dose. These fragments are rescued by fusion with cultured rodent cells, yielding subclones containing different portions of the human genome. This technique is described by Benham et al. (Genomics 4:509-517, 1989) and Cox et al., (Science 250:245-250, 1990). The random and independent nature of the subclones permits efficient mapping of any human genome marker. Human DNA isolated from a panel of 80-100 cell lines provides a mapping reagent for ordering EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids. In this approach, the frequency of breakage between markers is used to measure distance, allowing construction of fine resolution maps as has been done using conventional ESTs (Schuler et al., Science 274:540-546, 1996).

RH mapping has been used to generate a high-resolution whole genome radiation hybrid map of human chromosome 17q22-q25.3 across the genes for growth hormone (GH) and thymidine kinase (TK) (Foster et al., Genomics 33:185-192, 1996), the region surrounding the Gorlin syndrome gene (Obermayr et al., Eur. J. Hum. Genet. 4:242-245, 1996), 60 loci covering the entire short arm of chromosome 12 (Raeymaekers et al., Genomics 29:170-178, 1995), the region of human chromosome 22 containing the neurofibromatosis type 2 locus (Frazer et al., Genomics 14:574-584, 1992) and 13 loci on the long arm of chromosome 5 (Warrington et al., Genomics 11:701-708, 1991).

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EXAMPLE 44

Mapping of EST-related nucleic acids, positional segments of

EST-related nucleic acids or fragments of positional segments of

EST-related nucleic acids to Human Chromosomes using PCk techniques

EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids may be assigned to human chromosomes using PCR based methodologies. In such approaches, oligonucleotide primer pairs are designed from EST-related

nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids to minimize the chance of amplifying through an intron. Preferably, the oligonucleotide primers are 18-23 bp in length and are designed for PCR amplification. The creation of PCR primers from known sequences is well known to those with skill in the art. For a review of PCR technology see Erlich. in PCR Technology; Principles and Applications for DNA Amplification. 1992. W.H. Freeman and Co., New York.

The primers are used in polymerase chain reactions (PCR) to amplify templates from total human genomic DNA. PCR conditions are as follows: 60 ng of genomic DNA is used as a template for PCR with 80 ng of each oligonucleotide primer, 0.6 unit of Taq polymerase, and 1 μCu of a 32P-labeled deoxycytidine triphosphate. The PCR is performed in a microplate thermocycler (Techne) under the following conditions: 30 cycles of 94°C, 1.4 min; 55°C, 2 min; and 72°C, 2 min; with a final extension at 72°C for 10 min. The amplified products are analyzed on a 6% polyacrylamide sequencing gel and visualized by autoradiography. If the length of the resulting PCR product is identical to the distance between the ends of the primer sequences in the 5'EST from which the primers are derived, then the PCR reaction is repeated with DNA templates from two panels of human-rodent somatic cell hybrids, BIOS PCRable DNA (BIOS Corporation) and NIGMS Human-Rodent Somatic Cell Hybrid Mapping Panel Number 1 (NIGMS, Camden, NJ).

PCR is used to screen a series of somatic cell hybrid cell lines containing defined sets of human chromosomes for the presence of a given 5'EST. DNA is isolated from the somatic hybrids and used as starting templates for PCR reactions using the primer pairs from the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids. Only those somatic cell hybrids with chromosomes containing the human gene corresponding to the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids will yield an amplified fragment. The 5'ESTs are assigned to a chromosome by analysis of the segregation pattern of PCR products from the somatic hybrid DNA templates. The single human chromosome present in all cell hybrids that give rise to an amplified fragment is the chromosome containing that EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids. For a review of techniques and analysis of results from somatic cell gene mapping experiments. (See Ledbetter *et al.*, 30 Genomics 6:475-481 (1990)).

Alternatively, the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids may be mapped to individual chromosomes using FISH as described in Example 45 below.

EXAMPLE 45

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EST-related nucleic acids to Chromosomes Using

Fluorescence In Situ Hybridization

Fluorescence in situ hybridization allows the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids to be mapped to a particular location on a given chromosome. The chromosomes to be used for fluorescence in situ hybridization techniques may be obtained from a variety of sources including cell cultures, tissues, or whole blood.

In a preferred embodiment, chromosomal localization of EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids are obtained by FISH as described by Cherif et al. (Proc. Natl. Acad. Sci. U.S.A., 87:6639-6643, 1990). Metaphase chromosomes are prepared from phytohemagglutinin (PHA)-stimulated blood cell donors. PHA-stimulated lymphocytes from healthy males are cultured for 72 h in RPMI-1640 medium. For synchronization, methotrexate (10 µM) is added for 17 h, followed by addition of 5-bromodeoxyuridine (5-BrdU, 0.1 mM) for 6 h. Colcemid (1 µg/ml) is added for the last 15 min before harvesting the cells. 15 Cells are collected, washed in RPMI, incubated with a hypotonic solution of KCl (75 mM) at 37°C for 15 min and fixed in three changes of methanol:acetic acid (3:1). The cell suspension is dropped onto a glass slide and air dried. The EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids is labeled with biotin-16 dUTP by nick translation according to the manufacturer's instructions (Bethesda Research Laboratories, Bethesda, 20 MD), purified using a Sephadex G-50 column (Pharmacia, Upsala, Sweden) and precipitated. Just prior to hybridization, the DNA pellet is dissolved in hybridization buffer (50% formamide, 2 X SSC, 10% dextran sulfate, 1 mg/ml sonicated salmon sperm DNA, pH 7) and the probe is denatured at 70°C for 5-10 min.

Slides kept at -20°C are treated for 1 h at 37°C with RNase A (100 μg/ml), rinsed three times in 2 X SSC and dehydrated in an ethanol series. Chromosome preparations are denatured in 70% formamide, 2 X SSC for 2 min at 70°C, then dehydrated at 4°C. The slides are treated with proteinase K (10 μg/100 ml in 20 mM Tris-HCl, 2 mM CaCl₂) at 37°C for 8 min and dehydrated. The hybridization mixture containing the probe is placed on the slide, covered with a coverslip, sealed with rubber cement and incubated overnight in a humid chamber at 37°C. After hybridization and post-hybridization washes, the biotinylated probe is detected by avidin-FITC and amplified with additional layers of biotinylated goat anti-avidin and avidin-FITC. For chromosomal localization, fluorescent R-bands are obtained as previously described (Cherif et al., supra.). The slides are observed under a LEICA fluorescence microscope (DMRXA). Chromosomes are counterstained with propidium inclide and the fluorescent signal of the probe appears as two symmetrical vellow-green spots on both chromatids of the fluorescent R-band chromosome (red). Thus, a particular EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids may be

localized to a particular cytogenetic R-band on a given chromosome.

Once the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids have been assigned to particular chromosomes using the techniques described in Examples 42-44 above, they may be utilized to construct a high resolution map of the chromosomes on which they are located or to identify the chromosomes in a sample.

EXAMPLE 46

Use of EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids to Construct or Expand Chromosome Maps

10 Chromosome mapping involves assigning a given unique sequence to a particular chromosome as described above. Once the unique sequence has been mapped to a given chromosome, it is ordered relative to other unique sequences located on the same chromosome. One approach to chromosome mapping utilizes a series of yeast artificial chromosomes (YACs) bearing several thousand long inserts derived from the chromosomes of the organism from which the EST-related nucleic acids, positional 15 segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids are obtained. This approach is described in Ramaiah Nagaraja et al., Genome Research 7:210-222, March 1997. Briefly, in this approach each chromosome is broken into overlapping pieces which are inserted into the YAC vector. The YAC inserts are screened using PCR or other methods to determine whether they include the EST-related nucleic acids, positional segments of EST-related nucleic acids or 20 fragments of positional segments of EST-related nucleic acids whose position is to be determined. Once an insert has been found which includes the 5'EST, the insert can be analyzed by PCR or other methods to determine whether the insert also contains other sequences known to be on the chromosome or in the region from which the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids was derived. This process can be 25 repeated for each insert in the YAC library to determine the location of each of the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of ESTrelated nucleic acids relative to one another and to other known chromosomal markers. In this way, a high resolution map of the distribution of numerous unique markers along each of the organisms chromosomes may be obtained.

As described in Example 47 below EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids may also be used to identify genes associated with a particular phenotype, such as hereditary disease or drug response.

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3. Use of EST-related nucleic acids, positional segments of FST-related nucleic acids or fragments of positional segments of EST-related nucleic acids Gene Identification

EXAMPLE 47

This example illustrates an approach useful for the association of EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids with particular phenotypic characteristics. In this example, a particular EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids is used as a test probe to associate that EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids with a particular phenotypic characteristic.

EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids are mapped to a particular location on a human chromosome using techniques such as those described in Examples 41 and 42 or other techniques known in the art. A search of Mendelian Inheritance in Man (V. McKusick, Mendelian Inheritance in Man (available on line through Johns Hopkins University Welch Medical Library) reveals the region of the human chromosome which contains the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids to be a very gene rich region containing several known genes and several diseases or phenotypes for which genes have not been identified. The gene corresponding to this EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids thus becomes an immediate candidate for each of these genetic diseases.

Cells from patients with these diseases or phenotypes are isolated and expanded in culture. PCR primers from the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids are used to screen genomic DNA, mRNA or cDNA obtained from the patients. EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids that are not amplified in the patients can be positively associated with a particular disease by further analysis. Alternatively, the PCR analysis may yield fragments of different lengths when the samples are derived from an individual having the phenotype associated with the disease than when the sample is derived from a healthy individual, indicating that the gene containing the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids may be responsible for the genetic disease.

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VII. Use of EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids to Construct Vectors

The present EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids may also be used to construct secretion vectors capable of directing the secretion of the proteins encoded by genes therein. Such secretion vectors may facilitate the purification or enrichment of the proteins encoded by genes inserted therein by

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reducing the number of background proteins from which the desired protein must be purified or enriched. Exemplary secretion vectors are described in Example 48 below.

1. Construction of secretion vectors

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EXAMPLE 48

Construction of Secretion Vectors

The secretion vectors of the present invention include a promoter capable of directing gene expression in the host cell, tissue, or organism of interest. Such promoters include the Rous Sarcoma Virus promoter, the SV40 promoter, the human cytomegalovirus promoter, and other promoters familiar to those skilled in the art.

A signal sequence from one of the EST-related nucleic acids, positional segments of ESTrelated nucleic acids or fragments of positional segments of EST-related nucleic acids is operably linked to the promoter such that the mRNA transcribed from the promoter will direct the translation of the signal peptide. Preferably, the signal sequence is from one of the nucleic acids of SEQ ID NOs.24-811. The host cell, tissue, or organism may be any cell, tissue, or organism which recognizes the signal 15 peptide encoded by the signal sequence in the EST-related nucleic acids, positional segments of ESTrelated nucleic acids or fragments of positional segments of EST-related nucleic acids. Suitable hosts include mammalian cells, tissues or organisms, avian cells, tissues, or organisms, insect cells, tissues or organisms, or yeast.

In addition, the secretion vector contains cloning sites for inserting genes encoding the proteins 20 which are to be secreted. The cloning sites facilitate the cloning of the insert gene in frame with the signal sequence such that a fusion protein in which the signal peptide is fused to the protein encoded by the inserted gene is expressed from the mRNA transcribed from the promoter. The signal peptide directs the extracellular secretion of the fusion protein.

The secretion vector may be DNA or RNA and may integrate into the chromosome of the host, 25 be stably maintained as an extrachromosomal replicon in the host, be an artificial chromosome, or be transiently present in the host. Preferably, the secretion vector is maintained in multiple copies in each host cell. As used herein, multiple copies means at least 2, 5, 10, 20, 25, 50 or more than 50 copies per cell. In some embodiments, the multiple copies are maintained extrachromosomally. In other embodiments, the multiple copies result from amplification of a chromosomal sequence.

Many nucleic acid backbones suitable for use as secretion vectors are known to those skilled in the art, including retroviral vectors, SV40 vectors, Bovine Papilloma Virus vectors, yeast integrating plasmids, yeast episomal plasmids, yeast artificial chromosomes, human artificial chromosomes, P element vectors, baculovirus vectors, or bacterial plasmids capable of being transiently introduced into the host.

The secretion vector may also contain a polyA signal such that the polyA signal is located downstream of the gene inserted into the secretion vector.

After the gene encoding the protein for which secretion is desired is inserted into the secretion vector, the secretion vector is introduced into the host cell, tissue, or organism using calcium phosphate precipitation, DEAE-Dextran, electroporation, liposome-mediated transfection, viral particles or as naked DNA. The protein encoded by the inserted gene is then purified or enriched from the supernatant 5 using conventional techniques such as ammonium sulfate precipitation, immunoprecipitation, immunoaffinitychromatography, size exclusion chromatography, ion exchange chromatography, and HPLC. Alternatively, the secreted protein may be in a sufficiently enriched or pure state in the supernatant or growth media of the host to permit it to be used for its intended purpose without further enrichment.

The signal sequences may also be inserted into vectors designed for gene therapy. In such vectors, the signal sequence is operably linked to a promoter such that mRNA transcribed from the promoter encodes the signal peptide. A cloning site is located downstream of the signal sequence such that a gene encoding a protein whose secretion is desired may readily be inserted into the vector and fused to the signal sequence. The vector is introduced into an appropriate host cell. The protein 15 expressed from the promoter is secreted extracellularly, thereby producing a therapeutic effect.

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EXAMPLE 49

Fusion Vectors

The EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of 20 positional segments of EST-related nucleic acids may be used to construct fusion vectors for the expression of chimeric polypeptides. The chimeric polypeptides comprise a first polypeptide portion and a second polypeptide portion. In the fusion vectors of the present invention, nucleic acids encoding the first polypeptide portion and the second polypeptide portion are joined in frame with one another so as to generate a nucleic acid encoding the chimeric polypeptide. The nucleic acid encoding the chimeric 25 polypeptide is operably linked to a promoter which directs the expression of an mRNA encoding the chimeric polypeptide. The promoter may be in any of the expression vectors described herein including those described in Examples 21 and 48.

Preferably, the fusion vector is maintained in multiple copies in each host cell. In some embodiments, the multiple copies are maintained extrachromosomally. In other embodiments, the 30 multiple copies result from amplification of a chromosomal sequence.

The first polypeptide portion may comprise any of the polypeptides encoded by the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST related nucleic acids. In some embodiments, the first polypeptide portion may be one of the ESTrelated polypeptides, fragments of EST-related polypeptides, positional segments of EST-related 35 polypeptides, or fragments of positional segments of EST-related polypeptides.

The second polypeptide portion may comprise any polypeptide of interest. In some embodiments, the second polypeptide portion may comprise a polypeptide having a detectable

enzymatic activity such as green fluorescent protein or β galactosidase. Chimeric polypeptides in which the second polypeptide portion comprises a detectable polypeptide may be used to determine the intracellular localization of the first polypeptide portion. In such procedures, the fusion vector encoding the chimeric polypeptide is introduced into a host cell under conditions which facilitate the expression of 5 the chimeric polypeptide. Where appropriate, the cells are treated with a detection reagent which is visible under the microscope following a catalytic reaction with the detectable polypeptide and the cellular location of the detection reagent is determined. For example, if the polypeptide having a detectable enzymatic activity is β galactosidase, the cells may be treated with Xgal. Alternatively, where the detectable polypeptide is directly detectable without the addition of a detection reagent, the 10 intracellular location of the chimeric polypeptide is determined by performing microscopy under conditions in which the dectable polypeptide is visible. For example, if the detectable polypeptide is green fluorescent protein or a modified version thereof, microscopy is performed by exposing the host cells to light having an appropriate wavelength to cause the green fluorescent protein or modified version thereof to fluoresce.

Alternatively, the second polypeptide portion may comprise a polypeptide whose isolation, purification, or enrichment is desired. In such embodiments, the isolation, purification, or enrichment of the second polypeptide portion may be achieved by performing the immunoaffinity chromatography procedures described below using an immunoaffinity column having an antibody directed against the first polypeptide portion coupled thereto.

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The proteins encoded by the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids or the EST-related polypeptides, fragments of EST-related polypeptides, positional segments of EST-related polypeptides, or fragments of positional segments of EST-related polypeptides may also be used to generate antibodies as explained herein in order to identify the tissue type or cell species from which a sample is derived as 25 described in Example 50.

EXAMPLE 50

Identification of Tissue Types or Cell Species by Means of Labeled Tissue Specific Antibodies

Identification of specific tissues is accomplished by the visualization of tissue specific antigens by means of antibody preparations as described herein which are conjugated, directly or indirectly to a detectable marker. Selected labeled antibody species bind to their specific antigen binding partner in tissue sections, cell suspensions, or in extracts of soluble proteins from a dissue sample to provide a pattern for qualitative or semi-qualitative interpretation.

Antisera for these procedures must have a potency exceeding that of the native preparation, and for that reason, antibodies are concentrated to a mg/ml level by isolation of the gamma globulin fraction, for example, by ion-exchange chromatography or by ammonium sulfate fractionation. Also, to provide

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the most specific antisera, unwanted antibodies, for example to common proteins, must be removed from the gamma globulin fraction, for example by means of insoluble immunoabsorbents, before the antibodies are labeled with the marker. Either monoclonal or heterologous antisera is suitable for either procedure.

5 1. Immunohistochemical Techniques

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Purified, high-titer antibodies, prepared as described above, are conjugated to a detectable marker, as described, for example, by Fudenberg, H., Chap. 26 in: Basic 503 Clinical Immunology, 3rd Ed. Lange, Los Altos, California (1980) or Rose, et al., Chap. 12 in: Methods in Immunodiagnosis, 2d Ed. John Wiley and Sons, New York (1980).

A fluorescent marker, either fluorescein or rhodamine, is preferred, but antibodies can also be labeled with an enzyme that supports a color producing reaction with a substrate, such as horseradish peroxidase. Markers can be added to tissue-bound antibody in a second step, as described below. Alternatively, the specific antitissue antibodies can be labeled with ferritin or other electron dense particles, and localization of the ferritin coupled antigen-antibody complexes achieved by means of an 15 electron microscope. In yet another approach, the antibodies are radiolabeled, with, for example 125I, and detected by overlaying the antibody treated preparation with photographic emulsion.

Preparations to carry out the procedures can comprise monoclonal or polyclonal antibodies to a single protein or peptide identified as specific to a tissue type, for example, brain tissue, or antibody preparations to several antigenically distinct tissue specific antigens can be used in panels, independently 20 or in mixtures, as required.

Tissue sections and cell suspensions are prepared for immunohistochemical examination according to common histological techniques. Multiple cryostat sections (about 4 µm, unfixed) of the unknown tissue and known control, are mounted and each slide covered with different dilutions of the antibody preparation. Sections of known and unknown tissues should also be treated with preparations 25 to provide a positive control, a negative control, for example, pre-immune sera, and a control for nonspecific staining, for example, buffer.

Treated sections are incubated in a humid chamber for 30 min at room temperature, rinsed, then washed in buffer for 30-45 min. Excess fluid is blotted away, and the marker developed.

If the tissue specific antibody was not labeled in the first incubation, it can be labeled at this time 30 in a second antibody-antibody reaction, for example, by adding fluorescein- or enzyme-conjugated antibody against the immunoglobulin class of the antiserum-producing species, for example, fluorescein labeled antibody to mouse IgG. Such labeled sera are commercially available.

The antigen found in the tissues by the above precedure can be quantified by measuring the intensity of color or fluorescence on the tissue section, and calibrating that signal using appropriate 35 standards.

2. Identification of Tissue Specific Soluble Proteins

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The visualization of tissue specific proteins and identification of unknown tissues from that procedure is carried out using the labeled antibody reagents and detection strategy as described for immunohistochemistry; however the sample is prepared according to an electrophoretic technique to distribute the proteins extracted from the tissue in an orderly array on the basis of molecular weight for 5 detection.

A tissue sample is homogenized using a Virtis apparatus; cell suspensions are disrupted by Dounce homogenization or osmotic lysis, using detergents in either case as required to disrupt cell membranes, as is the practice in the art. Insoluble cell components such as nuclei, microsomes, and membrane fragments are removed by ultracentrifugation, and the soluble protein-containing fraction 10 concentrated if necessary and reserved for analysis.

A sample of the soluble protein solution is resolved into individual protein species by conventional SDS polyacrylamide electrophoresis as described, for example, by Davis, L. et al., Section 19-2 in: Basic Methods in Molecular Biology (P. Leder, ed), Elsevier, New York (1986), using a range of amounts of polyacrylamide in a set of gels to resolve the entire molecular weight range of proteins to 15 be detected in the sample. A size marker is run in parallel for purposes of estimating molecular weights of the constituent proteins. Sample size for analysis is a convenient volume of from 5 to 55 µl, and containing from about 1 to 100 µg protein. An aliquot of each of the resolved proteins is transferred by blotting to a nitrocellulose filter paper, a process that maintains the pattern of resolution. Multiple copies are prepared. The procedure, known as Western Blot Analysis, is well described in Davis, L. et al., 20 supra Section 19-3. One set of nitrocellulose blots is stained with Coomassie Blue dye to visualize the entire set of proteins for comparison with the antibody bound proteins. The remaining nitrocellulose filters are then incubated with a solution of one or more specific antisera to tissue specific proteins prepared as described in Examples 20 and 33. In this procedure, as in procedure A above, appropriate positive and negative sample and reagent controls are run.

In either procedure described above a detectable label can be attached to the primary tissue antigen-primary antibody complex according to various strategies and permutations thereof. In a straightforward approach, the primary specific antibody can be labeled; alternatively, the unlabeled complex can be bound by a labeled secondary anti-IgG antibody. In other approaches, either the primary or secondary antibody is conjugated to a biotin molecule, which can, in a subsequent step, bind an avidin 30 conjugated marker. According to yet another strategy, enzyme labeled or radioactive protein A, which has the property of binding to any IgG, is bound in a final step to either the primary or secondary antibody.

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EXAMPLE 51

Immunohistochemical Localization of Polypeptides

The antibodies prepared as described herein above may be utilized to determine the cellular location of a polypeptide. The polypeptide may be any of the polypeptides encoded by EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids or the polypeptide may be one of the EST-related polypeptides, fragments of EST-related polypeptides, positional segments of EST-related polypeptides, or fragments of positional segments of EST-related polypeptides. In some embodiments, the polypeptide may be a chimeric polypeptide such as those encoded by the fusion vectors of Example 49.

Cells expressing the polypeptide to be localized are applied to a microscope slide and fixed using any of the procedures typically employed in immunohistochemical localization techniques, including the methods described in *Current Protocols in Molecular Biology*, John Wiley and Sons, Inc. 1997. Following a washing step, the cells are contacted with the antibody. In some embodiments, the antibody is conjugated to a detectable marker as described above to facilitate detection. Alternatively, in some embodiments, after the cells have been contacted with an antibody to the polypeptide to be localized, a secondary antibody which has been conjugated to a detectable marker is placed in contact with the antibody against the polypeptide to be localized.

Thereafter, microscopy is performed under conditions suitable for visualizing the cellular location of the polypeptide.

The visualization of tissue specific antigen binding at levels above those seen in control tissues to one or more tissue specific antibodies, directed against the polypeptides encoded by EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids or antibodies against the EST-related polypeptides, fragments of EST-related polypeptides, positional segments of EST-related polypeptides, or fragments of positional segments of EST-related polypeptides, can identify tissues of unknown origin, for example, forensic samples, or differentiated tumor tissue that has metastasized to foreign bodily sites.

The antibodies described herein may also be used in the immunoaffinity chromatography techniques described below to isolate, purify or enrich the polypeptides encoded by the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids or to isolate, purify or enrich EST-related polypeptides, fragments of EST-related polypeptides, positional segments of EST-related polypeptides, or fragments of positional segments of EST-related polypeptides. The immunoaffinity chromatography techniques described below may also be used to isolate, purify or enrich polypeptides which have been linked to the polypeptides encoded by the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids or to isolate, purify or enrich polypeptides which have been linked to EST-related polypeptides, fragments of EST-related polypeptides, positional segments of EST-related polypeptides, or fragments of positional segments of EST-related polypeptides, or fragments of positional segments of EST-related polypeptides, positional segments of EST-related polypeptides, or fragments of positional segments of EST-related polypeptides, positional segments of EST-related polypeptides.

127 **EXAMPLE 52**

Immunoaffinity Chromatography

Antibodies prepared as described above are coupled to a support. Preferably, the antibodies are monoclonal antibodies, but polyclonal antibodies may also be used. The support may be any of those typically employed in immunoaffinity chromatography, including Sepharose CL-4B (Pharmacia, Piscataway, NJ), Sepharose CL-2B (Pharmacia, Piscataway, NJ), Affi-gel 10 (Biorad, Richmond, CA), or glass beads.

The antibodies may be coupled to the support using any of the coupling reagents typically used in immunoaffinity chromatography, including cyanogen bromide. After coupling the antibody to the support, the support is contacted with a sample which contains a target polypeptide whose isolation, purification or enrichment is desired. The target polypeptide may be a polypeptide encoded by the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related polypeptides may be one of the EST-related polypeptides, fragments of EST-related polypeptides, positional segments of EST-related polypeptides. The target polypeptides may also be polypeptides which have been linked to the polypeptides encoded by the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids or fragments of positional segments of EST-related polypeptides which have been linked to EST-related polypeptides, fragments of EST-related polypeptides, positional segments of EST-related polypeptides, or fragments of positional segments of EST-related polypeptides using the fusion vectors described above.

Preferably, the sample is placed in contact with the support for a sufficient amount of time and under appropriate conditions to allow at least 50% of the target polypeptide to specifically bind to the antibody coupled to the support.

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Thereafter, the support is washed with an appropriate wash solution to remove polypeptides which have non-specifically adhered to the support. The wash solution may be any of those typically employed in immunoaffinity chromatography, including PBS, Tris-lithium chloride buffer (0.1M lysine base and 0.5M lithium chloride, pH 8.0), Tris-hydrochloride buffer (0.05M Tris-hydrochloride, pH 8.0), or Tris/Triton/NaCl buffer (50mM Tris.cl, pH 8.0 or 9.0, 0.1% Triton X-100, and 0.5MNaCl).

After washing, the specifically bound target polypeptide is eluted from the support using the high pH or low pH elution solutions typically employed in immunoaffinity chromatography. In particular, the elution solutions may contain an eluant such as triethanolamine, diethylamine, calcium chioride, sodium thiocyanate, potasssium bromide, acetic acid, or glycine. In some embediments, the elution solution may also contain a detergent such as Triton X-100 or octyl-β-D-glucoside.

The EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids may also be used to clone sequences located upstream of the 5'ESTs which are capable of regulating gene expression, including promoter sequences, enhancer

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sequences, and other upstream sequences which influence transcription or translation levels. Once identified and cloned, these upstream regulatory sequences may be used in expression vectors designed to direct the expression of an inserted gene in a desired spatial, temporal, developmental, or quantitative fashion. Example 51 describes a method for cloning sequences upstream of the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids.

2. Identification of upstream sequences with promoting or regulatory activities

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EXAMPLE 53

Use of EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids to Clone Upstream Sequences from Genomic DNA

Sequences derived from EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids may be used to isolate the promoters of the corresponding genes using chromosome walking techniques. In one chromosome walking technique, which utilizes the GenomeWalkerTM kit available from Clontech, five complete genomic DNA samples are each digested with a different restriction enzyme which has a 6 base recognition site and leaves a blunt end. Following digestion, oligonucleotide adapters are ligated to each end of the resulting genomic DNA fragments.

For each of the five genomic DNA libraries, a first PCR reaction is performed according to the manufacturer's instructions using an outer adapter primer provided in the kit and an outer gene specific primer. The gene specific primer should be selected to be specific for 5' EST of interest and should have a melting temperature, length, and location in the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids which is consistent with its use in PCR reactions. Each first PCR reaction contains 5ng of genomic DNA, 5 μl of 10X Tth reaction buffer, 0.2 mM of each dNTP, 0.2 μM each of outer adapter primer and outer gene specific primer, 1.1 mM of Mg(OAc)₂, and 1 μl of the Tth polymerase 50X mix in a total volume of 50 μl. The reaction cycle for the first PCR reaction is as follows: 1 min at 94°C / 2 sec at 94°C, 3 min at 72°C (7 cycles) / 2 sec at 94°C, 3 min at 67°C (32 cycles) / 5 min at 67°C.

The product of the first PCR reaction is diluted and used as a template for a second PCR

reaction according to the manufacturer's instructions using a pair of nested primers which are located internally on the amplicon resulting from the first PCR reaction. For example, 5 µl of the reaction product of the first PCR reaction mixture may be diluted 180 times. Reactions are made in a 50 µl volume having a composition identical to that of the first PCR reaction except the nested primers are used. The first nested primer is specific for the adapter, and is provided with the GenomeWalkerTM

kit. The second nested primer is specific for the particular EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids for which the promoter is to be cloned and should have a melting temperature, length, and location in

the EST-related nucleic acids, positi nal segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids which is consistent with its use in PCR reactions.

The reaction parameters of the second PCR reaction are as follows: 1 min at 94°C / 2 sec at 94°C, 3 min at 72°C (6 cycles) / 2 sec at 94°C, 3 min at 67°C (25 cycles) / 5 min at 67°C. The product of the second PCR reaction is purified, cloned, and sequenced using standard techniques.

Alternatively, two or more human genomic DNA libraries can be constructed by using two or more restriction enzymes. The digested genomic DNA is cloned into vectors which can be converted into single stranded, circular, or linear DNA. A biotinylated oligonucleotide comprising at least 10, 12, 15, 18, 20, 23, 25, 27, 30, 35, 40, or 50 nucleotides from the EST-related nucleic acids, positional 10 segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids sequence is hybridized to the single stranded DNA. Hybrids between the biotinylated oligonucleotide and the single stranded DNA containing the EST-related nucleic acids, positional segments of ESTrelated nucleic acids or fragments of positional segments of EST-related nucleic acids are isolated as described above. Thereafter, the single stranded DNA containing the EST-related nucleic acids, 15 positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids is released from the beads and converted into double stranded DNA using a primer specific for the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids or a primer corresponding to a sequence included in the cloning vector. The resulting double stranded DNA is transformed into bacteria. cDNAs containing the 20 EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids are identified by colony PCR or colony hybridization.

Once the upstream genomic sequences have been cloned and sequenced as described above, prospective promoters and transcription start sites within the upstream sequences may be identified by comparing the sequences upstream of the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids with databases containing known transcription start sites, transcription factor binding sites, or promoter sequences.

In addition, promoters in the upstream sequences may be identified using promoter reporter vectors as described in Example 54.

EXAMPLE 54

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Identification of Promoters in Cloned Upstream Sequences

The genomic sequences upstream of the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids are cloned into a suitable promoter reporter vector, such as the pSEAP-Basic, pSEAP-Enhancer, pβ-gal-Basic, pβ-gal-B

protein. The sequences upstream of the EST-related nucleic acids, positional segments of EST-related nucleic acids are inserted into the cloning sites upstream of the reporter gene in both orientations and introduced into an appropriate host cell. The level of reporter protein is assayed and compared to the level obtained from a vector which lacks an insert in the cloning site. The presence of an elevated expression level in the vector containing the insert with respect to the control vector indicates the presence of a promoter in the insert. If necessary, the upstream sequences can be cloned into vectors which contain an enhancer for augmenting transcription levels from weak promoter sequences. A significant level of expression above that observed with the vector lacking an insert indicates that a promoter sequence is present in the inserted upstream sequence.

Appropriate host cells for the promoter reporter vectors may be chosen based on the results of the above described determination of expression patterns of the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids. For example, if the expression pattern analysis indicates that the mRNA corresponding to a particular EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids is expressed in fibroblasts, the promoter reporter vector may be introduced into a human fibroblast cell line.

Promoter sequences within the upstream genomic DNA may be further defined by constructing nested deletions in the upstream DNA using conventional techniques such as Exonuclease III digestion.

The resulting deletion fragments can be inserted into the promoter reporter vector to determine whether the deletion has reduced or obliterated promoter activity. In this way, the boundaries of the promoters may be defined. If desired, potential individual regulatory sites within the promoter may be identified using site directed mutagenesis or linker scanning to obliterate potential transcription factor binding sites within the promoter individually or in combination. The effects of these mutations on transcription levels may be determined by inserting the mutations into the cloning sites in the promoter reporter vectors.

EXAMPLE 55

Cloning and Identification of Promoters

30 Using the method described in Example 54 above with 5' ESTs, sequences upstream of several genes were obtained. Using the primer pairs GGG AAG ATG GAG ATA GTA TTG CCT G (SEQ ID NO:15) and CTG CCA TGT ACA TGA TAG AGA GAT TC (SEQ ID NO:16), the promoter having the internal designation P13412 (SEQ ID NO:17) was obtained.

TIsing the primer pairs GTA-CCA GGGG ACT GTG ACC ATT GC (SECTD NO:18) and CTG-35 TGA CCA TTG CTC CCA AGA GAG (SEQ ID NO:19), the promoter having the internal designation P15B4 (SEQ ID NO:20) was obtained. WO 99/53051 PCT/IB99/00712

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Using the primer pairs CTG GGA TGG AAG GCA CGG TA (SEQ ID NO:21) and GAG ACC ACA CAG CTA GAC AA (SEQ ID NO:22), the promoter having the internal designation P29B6 (SEQ ID NO:23) was obtained.

Figure 4 provides a schematic description of the promoters isolated and the way they are

seembled with the corresponding 5' tags. The upstream sequences were screened for the presence of
motifs resembling transcription factor binding sites or known transcription start sites using the computer
program MatInspector release 2.0, August 1996.

Figure 5 describes the transcription factor binding sites present in each of these promoters. The columns labeled matrice provides the name of the MatInspector matrix used. The column labeled position provides the 5' position of the promoter site. Numeration of the sequence starts from the transcription site as determined by matching the genomic sequence with the 5' EST sequence. The column labeled "orientation" indicates the DNA strand on which the site is found, with the + strand being the coding strand as determined by matching the genomic sequence with the sequence of the 5' EST. The column labeled "score" provides the MatInspector score found for this site. The column labeled "length" provides the length of the site in nucleotides. The column labeled "sequence" provides the sequence of the site found.

Bacterial clones containing plasmids containing the promoter sequences described above described above are presently stored in the inventor's laboratories under the internal identification numbers provided above. The inserts may be recovered from the deposited materials by growing an aliquot of the appropriate bacterial clone in the appropriate medium. The plasmid DNA can then be isolated using plasmid isolation procedures familiar to those skilled in the art such as alkaline lysis minipreps or large scale alkaline lysis plasmid isolation procedures. If desired the plasmid DNA may be further enriched by centrifugation on a cesium chloride gradient, size exclusion chromatography, or anion exchange chromatography. The plasmid DNA obtained using these procedures may then be manipulated using standard cloning techniques familiar to those skilled in the art. Alternatively, a PCR can be done with primers designed at both ends of the inserted EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids. The PCR product which corresponds to the EST-related nucleic acids, positional segments of EST-related nucleic acids can then be manipulated using standard cloning techniques familiar to those skilled in the art.

The promoters and other regulatory sequences located upstream of the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids may be used to design expression vectors capable of directing the expression of an inserted gene in a desired spatial, temporal, developmental, or quantifative manner. A promoter capable of directing the desired spatial, temporal, developmental, and quantitative patterns may be selected using the results of the expression analysis described above. For example, if a promoter which confers a high level of expression in muscle is desired, the promoter sequence upstream of EST-related nucleic acids,

positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids derived from an mRNA which are expressed at a high level in muscle, as determined by the methods above, may be used in the expression vector.

Preferably, the desired promoter is placed near multiple restriction sites to facilitate the cloning of the desired insert downstream of the promoter, such that the promoter is able to drive expression of the inserted gene. The promoter may be inserted in conventional nucleic acid backbones designed for extrachromosomal replication, integration into the host chromosomes or transient expression. Suitable backbones for the present expression vectors include retroviral backbones, backbones from eukaryotic episomes such as SV40 or Bovine Papilloma Virus, backbones from bacterial episomes, or artificial chromosomes.

Preferably, the expression vectors also include a polyA signal downstream of the multiple restriction sites for directing the polyadenylation of mRNA transcribed from the gene inserted into the expression vector.

Following the identification of promoter sequences, proteins which interact with the promoter may be identified as described in Example 56 below.

EXAMPLE 56

Identification of Proteins Which Interact with

Promoter Sequences, Upstream Regulatory Sequences, or mRNA

Sequences within the promoter region which are likely to bind transcription factors may be identified by homology to known transcription factor binding sites or through conventional mutagenesis or deletion analyses of reporter plasmids containing the promoter sequence. For example, deletions may be made in a reporter plasmid containing the promoter sequence of interest operably linked to an assayable reporter gene. The reporter plasmids carrying various deletions within the promoter region are transfected into an appropriate host cell and the effects of the deletions on expression levels is assessed. Transcription factor binding sites within the regions in which deletions reduce expression levels may be further localized using site directed mutagenesis, linker scanning analysis, or other techniques familiar to those skilled in the art.

Nucleic acids encoding proteins which interact with sequences in the promoter may be identified using one-hybrid systems such as those described in the manual accompanying the Matchmaker One-Hybrid System kit available from Clontech (Catalog No. K1603-1). Briefly, the Matchmaker One-hybrid system is used as follows. The target sequence for which it is desired to identify binding proteins is cloned upstream of a selectable reporter gene and integrated into the yeast general. Preferably, multiple copies of the target sequences are inserted into the reporter plasmid in tandem. A library comprised of fusions between cDNAs to be evaluated for the ability to bind to the promoter and the activation domain of a yeast transcription factor, such as GAL4, is transformed into the yeast strain containing the integrated reporter sequence. The yeast are plated on selective media to select cells

expressing the selectable marker linked to the promoter sequence. The colonies which grow on the selective media contain genes encoding proteins which bind the target sequence. The inserts in the genes encoding the fusion proteins are further characterized by sequencing. In addition, the inserts may be inserted into expression vectors or in vitro transcription vectors. Binding of the polypeptides encoded by 5 the inserts to the promoter DNA may be confirmed by techniques familiar to those skilled in the art, such as gel shift analysis or DNAse protection analysis.

VIII. Use of EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids in Gene Therapy

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The present invention also comprises the use of EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids in gene therapy strategies, including antisense and triple helix strategies as described in Examples 57 and 58 below. In antisense approaches, nucleic acid sequences complementary to an mRNA are hybridized to the mRNA intracellularly, thereby blocking the expression of the protein encoded by the mRNA. The 15 antisense sequences may prevent gene expression through a variety of mechanisms. For example, the antisense sequences may inhibit the ability of ribosomes to translate the mRNA. Alternatively, the antisense sequences may block transport of the mRNA from the nucleus to the cytoplasm, thereby limiting the amount of mRNA available for translation. Another mechanism through which antisense sequences may inhibit gene expression is by interfering with mRNA splicing. In yet another strategy, 20 the antisense nucleic acid may be incorporated in a ribozyme capable of specifically cleaving the target mRNA.

EXAMPLE 57

Preparation and Use of Antisense Oligonucleotides

25 The antisense nucleic acid molecules to be used in gene therapy may be either DNA or RNA sequences. They may comprise a sequence complementary to the sequence of the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of ESTrelated nucleic acids. The antisense nucleic acids should have a length and melting temperature sufficient to permit formation of an intracellular duplex with sufficient stability to inhibit the expression 30 of the mRNA in the duplex. Strategies for designing antisense nucleic acids suitable for use in gene therapy are disclosed in Green et al., Ann. Rev. Biochem. 55:569-597 (1986) and Izant and Weintraub, Cell 36:1007-1015 (1984).

In some strategies, antisense molecules are obtained from a molecule sequence encoding a protein by reversing the orientation of the coding region with respect to a promoter so as to transcribe the 35 opposite strand from that which is normally transcribed in the cell. The antisense molecules may be transcribed using in vitro transcription systems such as those which employ T7 or SP6 polymerase to

generate the transcript. Another approach involves transcription of the antisense nucleic acids *in vivo* by operably linking DNA containing the antisense sequence to a promoter in an expression vector.

Alternatively, oligonucleotides which are complementary to the strand normally transcribed in the cell may be synthesized *in vitro*. Thus, the antisense nucleic acids are complementary to the corresponding mRNA and are capable of hybridizing to the mRNA to create a duplex. In some embodiments, the antisense sequences may contain modified sugar phosphate backbones to increase stability and make them less sensitive to RNase activity. Examples of modifications suitable for use in antisense strategies are described by Rossi *et al.*, *Pharmacol. Ther.* **50(2)**:245-254, (1991).

Various types of antisense oligonucleotides complementary to the sequence of the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids may be used. In one preferred embodiment, stable and semi-stable antisense oligonucleotides described in International Application No. PCT WO94/23026 are used. In these molecules, the 3' end or both the 3' and 5' ends are engaged in intramolecular hydrogen bonding between complementary base pairs. These molecules are better able to withstand exonuclease attacks and exhibit increased stability compared to conventional antisense oligonucleotides.

In another preferred embodiment, the antisense oligodeoxynucleotides against herpes simplex virus types 1 and 2 described in International Application No. WO 95/04141 are used.

In yet another preferred embodiment, the covalently cross-linked antisense oligonucleotides described in International Application No. WO 96/31523 are used. These double- or single-stranded oligonucleotides comprise one or more, respectively, inter- or intra-oligonucleotide covalent cross-linkages, wherein the linkage consists of an amide bond between a primary amine group of one strand and a carboxyl group of the other strand or of the same strand, respectively, the primary amine group being directly substituted in the 2' position of the strand nucleotide monosaccharide ring, and the carboxyl group being carried by an aliphatic spacer group substituted on a nucleotide or nucleotide analog of the other strand or the same strand, respectively.

The antisense oligodeoxynucleotides and oligonucleotides disclosed in International Application No. WO 92/18522 may also be used. These molecules are stable to degradation and contain at least one transcription control recognition sequence which binds to control proteins and are effective as decoys therefor. These molecules may contain "hairpin" structures, "dumbbell" structures, "modified dumbbell" structures, "cross-linked" decoy structures and "loop" structures.

In another preferred embodiment, the cyclic double-stranded oligonucleotides described in European Patent Application No. 0 572 287 A2. These ligated oligonucleotide "dumbbells" contain the binding site for a transcription factor and inhibit expression of the gene under control of the transcription factor by sequestering the factor.

Use of the closed antisense oligonucleotides disclosed in International Application No. WO 92/19732 is also contemplated. Because these molecules have no free ends, they are more resistant to

degradation by exonucleases than are conventional oligonucleotides. These oligonucleotides may be multifunctional, interacting with several regions which are not adjacent to the target mRNA.

The appropriate level of antisense nucleic acids required to inhibit gene expression may be determined using *in vitro* expression analysis. The antisense molecule may be introduced into the cells by diffusion, injection, infection or transfection using procedures known in the art. For example, the antisense nucleic acids can be introduced into the body as a bare or naked oligonucleotide, oligonucleotide encapsulated in lipid, oligonucleotide sequence encapsidated by viral protein, or as an oligonucleotide operably linked to a promoter contained in an expression vector. The expression vector may be any of a variety of expression vectors known in the art, including retroviral or viral vectors, vectors capable of extrachromosomal replication, or integrating vectors. The vectors may be DNA or RNA.

The antisense molecules are introduced onto cell samples at a number of different concentrations preferably between $1 \times 10^{-10} M$ to $1 \times 10^{-4} M$. Once the minimum concentration that can adequately control gene expression is identified, the optimized dose is translated into a dosage suitable for use *in vivo*. For example, an inhibiting concentration in culture of 1×10^{-7} translates into a dose of approximately 0.6 mg/kg bodyweight. Levels of oligonucleotide approaching 100 mg/kg bodyweight or higher may be possible after testing the toxicity of the oligonucleotide in laboratory animals. It is additionally contemplated that cells from the vertebrate are removed, treated with the antisense oligonucleotide, and reintroduced into the vertebrate.

It is further contemplated that the antisense oligonucleotide sequence is incorporated into a ribozyme sequence to enable the antisense to specifically bind and cleave its target mRNA. For technical applications of ribozyme and antisense oligonucleotides see Rossi *et al.*, *supra*.

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In a preferred application of this invention, the polypeptide encoded by the gene is first identified, so that the effectiveness of antisense inhibition on translation can be monitored using techniques that include but are not limited to antibody-mediated tests such as RIAs and ELISA, functional assays, or radiolabeling.

The EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids may also be used in gene therapy approaches based on intracellular triple helix formation. Triple helix oligonucleotides are used to inhibit transcription from a genome. They are particularly useful for studying alterations in cell activity as it is associated with a particular gene. The EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids of the present invention or, more preferably, a portion of those sequences, can be used to inhibit gene expression in individuals having diseases associated with expression of a particular gene. Similarly, the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids or fragments of EST-related nucleic acids or fragments of EST-related nucleic ac

However, homopyrimidine sequences can also inhibit gene expression. Such homopyrimidine oligonucleotides bind to the major groove at homopurine:homopyrimidine sequences. Thus, both types of sequences from the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids are contemplated within the scope of this invention.

EXAMPLE 58

Preparation and use of Triple Helix Probes

The sequences of the EST-related nucleic acids, positional segments of EST-related nucleic acids are scanned to identify 10-mer to 20-mer homopyrimidine or homopurine stretches which could be used in triple-helix based strategies for inhibiting gene expression. Following identification of candidate homopyrimidine or homopurine stretches, their efficiency in inhibiting gene expression is assessed by introducing varying amounts of oligonucleotides containing the candidate sequences into tissue culture cells which normally express the target gene. The oligonucleotides may be prepared on an oligonucleotide synthesizer or they may be purchased commercially from a company specializing in custom oligonucleotide synthesis, such as GENSET, Paris, France.

The oligonucleotides may be introduced into the cells using a variety of methods known to those skilled in the art, including but not limited to calcium phosphate precipitation, DEAE-Dextran,

20 electroporation, liposome-mediated transfection or native uptake.

Treated cells are monitored for altered cell function or reduced gene expression using techniques such as Northern blotting, RNase protection assays, or PCR based strategies to monitor the transcription levels of the target gene in cells which have been treated with the oligonucleotide. The cell functions to be monitored are predicted based upon the homologies of the target genes corresponding to the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids from which the oligonucleotide were derived with known gene sequences that have been associated with a particular function. The cell functions can also be predicted based on the presence of abnormal physiologies within cells derived from individuals with a particular inherited disease, particularly when the EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids are associated with the disease using techniques described herein.

The oligonucleotides which are effective in inhibiting gene expression in tissue culture cells may then be introduced *in vivo* using the techniques described above and in Example 56 at a dosage calculated based on the *in vitro* results, as described in Example 57.

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In some embodiments, the natural (beta) anomers of the oligonucleotide units can be replaced with alpha anomers to render the oligonucleotide more resistant to nucleases. Further, an intercalating agent such as ethidium bromide, or the like, can be attached to the 3' end of the alpha oligonucleotide to

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stabilize the triple helix. For information on the generation of oligonucleotides suitable for triple helix formation see Griffin et al. (Science 245:967-971 (1989)).

EXAMPLE 59

Use of EST-related nucleic acids, positional segments of

EST-related nucleic acids or fragments of positional segments of

EST-related nucleic acids to express an Encoded Protein in a Host Organism

The EST-related nucleic acids, positional segments of EST-related nucleic acids or fragments of positional segments of EST-related nucleic acids may also be used to express an encoded protein or polypeptide in a host organism to produce a beneficial effect. In addition, nucleic acids encoding the EST-related polypeptides, positional segments of EST-related polypeptides or fragments of positional segments of EST-related polypeptides may be used to express the encoded protein or polypeptide in a host organism to produce a beneficial effect.

In such procedures, the encoded protein or polypeptide may be transiently expressed in the host organism or stably expressed in the host organism. The encoded protein or polypeptide may have any of the activities described above. The encoded protein or polypeptide may be a protein or polypeptide which the host organism lacks or, alternatively, the encoded protein may augment the existing levels of the protein in the host organism.

In some embodiments in which the protein or polypeptide is secreted, nucleic acids encoding the full length protein (i.e. the signal peptide and the mature protein), or nucleic acids encoding only the mature protein (i.e. the protein generated when the signal peptide is cleaved off) is introduced into the host organism.

The nucleic acids encoding the proteins or polypeptides may be introduced into the host organism using a variety of techniques known to those of skill in the art. For example, the extended cDNA may be injected into the host organism as naked DNA such that the encoded protein is expressed in the host organism, thereby producing a beneficial effect.

Alternatively, the nucleic acids encoding the protein or polypeptide may be cloned into an expression vector downstream of a promoter which is active in the host organism. The expression vector may be any of the expression vectors designed for use in gene therapy, including viral or retroviral vectors. The expression vector may be directly introduced into the host organism such that the encoded protein is expressed in the host organism to produce a beneficial effect. In another approach, the expression vector may be introduced into cells *in vitro*. Cells containing the expression vector are thereafter selected and introduced into the host organism, where they express the expression protein or polypeptide to produce a beneficial effect.

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The short core hydrophobic region (h) of signal peptides encoded by the sequences of SEQ ID NOs. 24-728 and 766-792 may also be used as a carrier to import a peptide or a protein of interest, socalled cargo, into tissue culture cells (Lin et al., J. Biol. Chem., 270: 14225-14258 (1995); Du et al., J. Peptide Res., 51: 235-243 (1998); Rojas et al., Nature Biotech., 16: 370-375 (1998)).

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When cell permeable peptides of limited size (approximately up to 25 amino acids) are to be translocated across cell membrane, chemical synthesis may be used in order to add the h region to either the C-terminus or the N-terminus to the cargo peptide of interest. Alternatively, when longer peptides or proteins are to be imported into cells, nucleic acids can be genetically engineered, using techniques familiar to those skilled in the art, in order to link the extended cDNA sequence encoding the h region to 10 the 5' or the 3' end of a DNA sequence coding for a cargo polypeptide. Such genetically engineered nucleic acids are then translated either in vitro or in vivo after transfection into appropriate cells, using conventional techniques to produce the resulting cell permeable polypeptide. Suitable hosts cells are then simply incubated with the cell permeable polypeptide which is then translocated across the membrane.

This method may be applied to study diverse intracellular functions and cellular processes. For 15 instance, it has been used to probe functionally relevant domains of intracellular proteins and to examine protein-protein interactions involved in signal transduction pathways (Lin et al., supra; Lin et al., J. Biol. Chem., 271: 5305-5308 (1996); Rojas et al., J. Biol. Chem., 271: 27456-27461 (1996); Liu et al., Proc. Natl. Acad. Sci. USA, 93: 11819-11824 (1996); Rojas et al., Bioch. Biophys. Res. Commun., 234: 675-20 680 (1997)).

Such techniques may be used in cellular therapy to import proteins producing therapeutic effects. For instance, cells isolated from a patient may be treated with imported therapeutic proteins and then re-introduced into the host organism.

Alternatively, the h region of signal peptides of the present invention could be used in 25 combination with a nuclear localization signal to deliver nucleic acids into cell nucleus. Such oligonucleotides may be antisense oligonucleotides or oligonucleotides designed to form triple helixes, as describedabove, in order to inhibit processing and maturation of a target cellular RNA.

EXAMPLE 61

Computer Embodiments

As used herein the term "nucleic acid codes of SEQ ID NOs. 24-811 and 1600-1622" encompasses the nucleotide sequences of SEQ ID NOs. 24-811 and 1600-1622, fragments of SEQ ID NOs. 24-811 and 1600-1622, nucleotide sequences homologous to SEQ ID NOs. 24-811 and 1600-1622 or homotogous to fragments of SEO ID NOs, 24-814 and 1600-1622, and sequences 35 complementary to all of the preceding sequences. The fragments include portions of SEQ ID NOs. 24-811 and 1600-1622 comprising at least 10, 15, 20, 25, 30, 35, 40, 50, 75, 100, 150, 200, 300, 400, or 500 consecutive nucleotides of SEQ ID NOs. 24-811 and 1600-1622. Preferably, the fragments are novel

fragments. Preferably the fragments include polynucleotides described in Table II, polynucleotides described in Table III, polynucleotides described in Table IV or portions thereof comprising at least 10, 15, 20, 25, 30, 35, 40, 50, 75, 100, 150, 200, 300, 400, or 500 consecutive nucleotides of the polynucleotides described in Tables II, III, or IV. Homologous sequences and fragments of SEQ ID NOs. 24-811 and 1600-1622 refer to a sequence having at least 99%, 98%, 97%, 96%, 95%, 90%, 85%, 80%, or 75% homology to these sequences. Homology may be determined using any of the computer programs and parameters described in Example 18, including BLAST2N with the default parameters or with any modified parameters. Homologous sequences also include RNA sequences in which uridines replace the thymines in the nucleic acid codes of SEQ ID NOs. 24-811 and 1600-1622. The

homologous sequences may be obtained using any of the procedures described herein or may result from the correction of a sequencing error as described above. Preferably the homologous sequences and fragments of SEQ ID NOs. 24-811 and 1600-1622 include polynucleotides described in Table II, polynucleotides described in Table IV or portions thereof comprising at least 10, 15, 20, 25, 30, 35, 40, 50, 75, 100, 150, 200, 300, 400, or 500 consecutive
 nucleotides of the polynucleotides described in Tables II, III, or IV. It will be appreciated that the nucleic acid codes of SEQ ID NOs. 24-811 and 1600-1622 can be represented in the traditional single character format (See the inside back cover of Styer, Lubert. *Biochemistry*, 3rd edition. W. H Freeman &

Co., New York.) or in any other format which records the identity of the nucleotides in a sequence.

As used herein the term "polypeptide codes of SEQ ID NOS. 812-1599" encompasses the 20 polypeptide sequence of SEQ ID NOs. 812-1599 which are encoded by the 5' EST s of SEQ ID NOs. 24-811 and 1600-1622, polypeptide sequences homologous to the polypeptides of SEQ ID NOS. 812-1599, or fragments of any of the preceding sequences. Homologous polypeptide sequences refer to a polypeptide sequence having at least 99%, 98%, 97%, 96%, 95%, 90%, 85%, 80%, 75% homology to one of the polypeptide sequences of SEQ ID NOS. 812-1599. Homology may be determined using any 25 of the computer programs and parameters described herein, including FASTA with the default parameters or with any modified parameters. The homologous sequences may be obtained using any of the procedures described herein or may result from the correction of a sequencing error as described above. The polypeptide fragments comprise at least 5, 10, 15, 20, 25, 30, 35, 40, 50, 75, 100, or 150 consecutive amino acids of the polypeptides of SEQ ID NOS. 812-1599. Preferably, the fragments are 30 novel fragments. Preferably, the fragments include polypeptides encoded by the polynucleotides described in Table II, or portions thereof comprising at least 5, 10, 15, 20, 25, 30, 35, 40, 50, 75, 100, or 150 consecutive amino acids of the polypeptides encoded by the polynucleotides described in Table II. It will be appreciated that the polypeptide codes of the SEQ ID NOS. 812-1599 can be represented in the traditional single character format or three letter format (See the inside back cover of Starrier, Libera 35 Biochemistry, 3rd edition. W. H Freeman & Co., New York.) or in any other format which relates the

identity of the polypeptides in a sequence.

It will be appreciated by those skilled in the art that the nucleic acid codes of SEQ ID NOs. 24-811 and 1600-1622 and polypeptide codes of SEQ ID NOS. 812-1599 can be stored, recorded, and manipulated on any medium which can be read and accessed by a computer. As used herein, the words "recorded" and "stored" refer to a process for storing information on a computer medium. A skilled artisan can readily adopt any of the presently known methods for recording information on a computer readable medium to generate manufactures comprising one or more of the nucleic acid codes of SEQ ID NOs. 24-811 and 1600-1622, one or more of the polypeptide codes of SEQ ID NOS. 812-1599.

Another aspect of the present invention is a computer readable medium having recorded thereon at least 2, 5, 10, 15, 20, 25, 30, or 50 nucleic acid codes of SEQ ID NOs. 24-811 and 1600-1622. Another aspect of the present invention is a computer readable medium having recorded thereon at least 2, 5, 10, 15, 20, 25, 30, or 50 polypeptide codes of SEQ ID NOS. 812-1599.

Computer readable media include magnetically readable media, optically readable media, electronically readable media and magnetic/optical media. For example, the computer readable media may be a hard disk, a floppy disk, a magnetic tape, CD-ROM, Digital Versatile Disk (DVD), Random

15 Access Memory (RAM), or Read Only Memory (ROM) as well as other types of other media known to those skilled in the art.

Embodiments of the present invention include systems, particularly computer systems which store and manipulate the sequence information described herein. One example of a computer system 100 is illustrated in block diagram form in Figure 6. As used herein, "a computer system" refers to the hardware components, software components, and data storage components used to analyze the nucleotide sequences of the nucleic acid codes of SEQ ID NOs. 24-811 and 1600-1622, or the amino acid sequences of the polypeptide codes of SEQ ID NOS. 812-1599. In one embodiment, the computer system 100 is a Sun Enterprise 1000 server (Sun Microsystems, Palo Alto, CA). The computer system 100 preferably includes a processor for processing, accessing and manipulating the sequence data. The 25 processor 105 can be any well-known type of central processing unit, such as the Pentium III from Intel Corporation, or similar processor from Sun, Motorola, Compaq or International Business Machines.

Preferably, the computer system 100 is a general purpose system that comprises the processor 105 and one or more internal data storage components 110 for storing data, and one or more data retrieving devices for retrieving the data stored on the data storage components. A skilled artisan can readily appreciate that any one of the currently available computer systems are suitable.

In one particular embodiment, the computer system 100 includes a processor 105 connected to a bus which is connected to a main memory 115 (preferably implemented as RAM) and one or more internal data storage devices 110. such as a hard drive and/or other computer readable media having data recorded thereon. In some embodiments, the computer system 100 further includes one or more data retrieving device 118 for reading the data stored on the internal data storage devices 110.

The data retrieving device 118 may represent, for example, a floppy disk drive, a compact disk drive, a magnetic tape drive, etc. In some embodiments, the internal data storage device 110 is a

removable computer readable medium such as a floppy disk, a compact disk, a magnetic tape, etc. containing control logic and/or data recorded thereon. The computer system 100 may advantageously include or be programmed by appropriate software for reading the control logic and/or the data from the data storage component once inserted in the data retrieving device.

The computer system 100 includes a display 120 which is used to display output to a computer user. It should also be noted that the computer system 100 can be linked to other computer systems 125a-c in a network or wide area network to provide centralized access to the computer system 100.

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Software for accessing and processing the nucleotide sequences of the nucleic acid codes of SEQ ID NOs. 24-811 and 1600-1622, or the amino acid sequences of the polypeptide codes of SEQ ID NOS. 812-1599 (such as search tools, compare tools, and modeling tools etc.) may reside in main memory 115 during execution.

In some embodiments, the computer system 100 may further comprise a sequence comparer for comparing the above-described nucleic acid codes of SEQ ID NOs. 24-811 and 1600-1622 or polypeptide codes of SEQ ID NOS. 812-1599 stored on a computer readable medium to reference nucleotide or polypeptide sequences stored on a computer readable medium. A "sequence comparer" refers to one or more programs which are implemented on the computer system 100 to compare a nucleotide or polypeptide sequence with other nucleotide or polypeptide sequences and/or compounds including but not limited to peptides, peptidomimetics, and chemicals stored within the data storage means. For example, the sequence comparer may compare the nucleotide sequences of the nucleic acid codes of SEQ ID NOs. 24-811 and 1600-1622, or the amino acid sequences of the polypeptide codes of SEQ ID NOS. 812-1599 stored on a computer readable medium to reference sequences stored on a computer readable medium to identify homologies, motifs implicated in biological function, or structural motifs. The various sequence comparer programs identified elsewhere in this patent specification are particularly contemplated for use in this aspect of the invention.

Figure 7 is a flow diagram illustrating one embodiment of a process 200 for comparing a new nucleotide or protein sequence with a database of sequences in order to determine the homology levels between the new sequence and the sequences in the database. The database of sequences can be a private database stored within the computer system 100, or a public database such as GENBANK, PIR OR SWISSPROT that is available through the Internet.

The process 200 begins at a start state 201 and then moves to a state 202 wherein the new sequence to be compared is stored to a memory in a computer system 100. As discussed above, the memory could be any type of memory, including RAM or an internal storage device.

The process 200 then moves to a state 204 wherein a database of sequences is opened for analysis and comparison. The process 200 then moves to a state 206 wherein the first sequence stored in the database is read into a memory on the computer. A comparison is then performed at a state 210 to determine if the first sequence is the same as the second sequence. It is important to note that this step is not limited to performing an exact comparison between the new sequence and the first sequence in the

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database. Well-known methods are known to those of skill in the art for comparing two nucleotide or protein sequences, even if they are not identical. For example, gaps can be introduced into one sequence in order to raise the homology level between the two tested sequences. The parameters that control whether gaps or other features are introduced into a sequence during comparison are normally entered by 5 the user of the computer system.

Once a comparison of the two sequences has been performed at the state 210, a determination is made at a decision state 210 whether the two sequences are the same. Of course, the term "same" is not limited to sequences that are absolutely identical. Sequences that are within the homology parameters entered by the user will be marked as "same" in the process 200.

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If a determination is made that the two sequences are the same, the process 200 moves to a state 214 wherein the name of the sequence from the database is displayed to the user. This state notifies the user that the sequence with the displayed name fulfills the homology constraints that were entered. Once the name of the stored sequence is displayed to the user, the process 200 moves to a decision state 218 wherein a determination is made whether more sequences exist in the database. If no more sequences 15 exist in the database, then the process 200 terminates at an end state 220. However, if more sequences do exist in the database, then the process 200 moves to a state 224 wherein a pointer is moved to the next sequence in the database so that it can be compared to the new sequence. In this manner, the new sequence is aligned and compared with every sequence in the database.

It should be noted that if a determination had been made at the decision state 212 that the 20 sequences were not homologous, then the process 200 would move immediately to the decision state 218 in order to determine if any other sequences were available in the database for comparison.

Accordingly, one aspect of the present invention is a computer system comprising a processor, a data storage device having stored thereon a nucleic acid code of SEQ ID NOs. 24-811 and 1600-1622 or a polypeptide code of SEQ ID NOS. 812-1599, a data storage device having 25 retrievably stored thereon reference nucleotide sequences or polypeptide sequences to be compared to the nucleic acid code of SEQ ID NOs. 24-811 and 1600-1622 or polypeptide code of SEQ ID NOS. 812-1599 and a sequence comparer for conducting the comparison. The sequence comparer may indicate a homology level between the sequences compared or identify structural motifs in the above described nucleic acid code of SEQ ID NOs. 24-811 and 1600-1622 and polypeptide codes of 30 SEQ ID NOS. 812-1599 or it may identify structural motifs in sequences which are compared to these nucleic acid codes and polypeptide codes. In some embodiments, the data storage device may have stored thereon the sequences of at least 2, 5, 10, 15, 20, 25, 30, or 50 of the nucleic acid codes of ©EQ ID NOs. 24-811 and 1600-1622 or polypeptide codes of SEQ ID NOS. 812-1599.

Another aspect of the present invention is a method for determining the level of homology 35 between a nucleic acid code of SEQ ID NOs. 24-811 and 1600-1622 and a reference nucleotide sequence, comprising the steps of reading the nucleic acid code and the reference nucleotide sequence through the use of a computer program which determines homology levels and determining homology

between the nucleic acid code and the reference nucleotide sequence with the computer program. The computer program may be any of a number of computer programs for determining homology levels, including those specifically enumerated herein, including BLAST2N with the default parameters or with any modified parameters. The method may be implemented using the computer systems described 5 above. The method may also be performed by reading 2, 5, 10, 15, 20, 25, 30, or 50 of the above described nucleic acid codes of SEQ ID NOs. 24-811 and 1600-1622 through use of the computer program and determining homology between the nucleic acid codes and reference nucleotide sequences.

Figure 8 is a flow diagram illustrating one embodiment of a process 250 in a computer for 10 determining whether two sequences are homologous. The process 250 begins at a start state 252 and then moves to a state 254 wherein a first sequence to be compared is stored to a memory. The second sequence to be compared is then stored to a memory at a state 256. The process 250 then moves to a state 260 wherein the first character in the first sequence is read and then to a state 262 wherein the first character of the second sequence is read. It should be understood that if the 15 sequence is a nucleotide sequence, then the character would normally be either A, T, C, G or U. If the sequence is a protein sequence, then it should be in the single letter amino acid code so that the first and sequence sequences can be easily compared.

A determination is then made at a decision state 264 whether the two characters are the same. If they are the same, then the process 250 moves to a state 268 wherein the next characters in the first 20 and second sequences are read. A determination is then made whether the next characters are the same. If they are, then the process 250 continues this loop until two characters are not the same. If a determination is made that the next two characters are not the same, the process 250 moves to a decision state 274 to determine whether there are any more characters either sequence to read.

If there aren't any more characters to read, then the process 250 moves to a state 276 wherein 25 the level of homology between the first and second sequences is displayed to the user. The level of homology is determined by calculating the proportion of characters between the sequences that were the same out of the total number of sequences in the first sequence. Thus, if every character in a first 100 nucleotide sequence aligned with a every character in a second sequence, the homology level would be 100%.

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Alternatively, the computer program may be a computer program which compares the nucleotide sequences of the nucleic acid codes of the present invention, to reference nucleotide sequences in order to determine whether the nucleic acid code of SEQ ID NOs. 24-811 and 1600-1622 differs from a reference nucleic acid sequence at one or more positions. Optionally such a program records the length and identity of inserted, deleted or substituted nucleatides with respect to the sequence 35 of either the reference polynucleotide or the nucleic acid code of SEQ ID NOs. 24-811 and 1600-1622. In one embodiment, the computer program may be a program which determines whether the nucleotide sequences of the nucleic acid codes of SEQ ID NOs. 24-811 and 1600-1622 contain a biallelic marker

or single nucleotide polymorphism (SNP) with respect to a reference nucleotide sequence. This single nucleotide polymorphism may comprise a single base substitution, insertion, or deletion, while this biallelic marker may comprise abour one to ten consecutive bases substituted, inserted or deleted.

Another aspect of the present invention is a method for determining the level of homology 5 between a polypeptide code of SEQ ID NOS. 812-1599 and a reference polypeptide sequence, comprising the steps of reading the polypeptide code of SEQ ID NOS. 812-1599 and the reference polypeptide sequence through use of a computer program which determines homology levels and determining homology between the polypeptide code and the reference polypeptide sequence using the computer program.

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Accordingly, another aspect of the present invention is a method for determining whether a nucleic acid code of SEQ ID NOs. 24-811 and 1600-1622 differs at one or more nucleotides from a reference nucleotide sequence comprising the steps of reading the nucleic acid code and the reference nucleotide sequence through use of a computer program which identifies differences between nucleic acid sequences and identifying differences between the nucleic acid code and the reference nucleotide 15 sequence with the computer program. In some embodiments, the computer program is a program which identifies single nucleotide polymorphisms. The method may be implemented by the computer systems described above and the method illustrated in Figure 8. The method may also be performed by reading at least 2, 5, 10, 15, 20, 25, 30, or 50 of the nucleic acid codes of SEQ ID NOs. 24-811 and 1600-1622 and the reference nucleotide sequences through the use of the computer program and identifying differences 20 between the nucleic acid codes and the reference nucleotide sequences with the computer program.

In other embodiments the computer based system may further comprise an identifier for identifying features within the nucleotide sequences of the nucleic acid codes of SEQ ID NOs. 24-811 and 1600-1622 or the amino acid sequences of the polypeptide codes of SEQ ID NOS. 812-1599.

An "identifier" refers to one or more programs which identifies certain features within the 25 above-described nucleotide sequences of the nucleic acid codes of SEQ ID NOs. 24-811 and 1600-1622 or the amino acid sequences of the polypeptide codes of SEQ ID NOS. 812-1599. In one embodiment, the identifier may comprise a program which identifies an open reading frame in the cDNAs codes of SEQ ID NOs. 24-811 and 1600-1622.

Figure 9 is a flow diagram illustrating one embodiment of an identifier process 300 for 30 detecting the presence of a feature in a sequence. The process 300 begins at a start state 302 and then moves to a state 304 wherein a first sequence that is to be checked for features is stored to a memory 115 in the computer system 100. The process 300 then moves to a state 306 wherein a database of sequence features is opened. Such a database would include a list of each feature's attributes along with the name of the feature. For example, a feature name could be "Initiation Codon" and the 35 attribute would be "ATG". Another example would be the feature name "TAATAA Box" and the feature attribute would be "TAATAA". An example of such a database is produced by the University of Wisconsin Genetics Computer Group (www.gcg.com).

Once the database of features is opened at the state 306, the process 300 moves to a state 308 wherein the first feature is read from the database. A comparison of the attribute of the first feature with the first sequence is then made at a state 310. A determination is then made at a decision state 316 whether the attribute of the feature was found in the first sequence. If the attribute was found, 5 then the process 300 moves to a state 318 wherein the name of the found feature is displayed to the user.

The process 300 then moves to a decision state 320 wherein a determination is made whether move features exist in the database. If no more features do exist, then the process 300 terminates at an end state 324. However, if more features do exist in the database, then the process 300 reads the next sequence feature at a state 326 and loops back to the state 310 wherein the attribute of the next feature is compared against the first sequence.

It should be noted, that if the feature attribute is not found in the first sequence at the decision state 316, the process 300 moves directly to the decision state 320 in order to determine if any more features exist in the database.

In another embodiment, the identifier may comprise a molecular modeling program which 15 determines the 3-dimensional structure of the polypeptides codes of SEQ ID NOS. 812-1599. In some embodiments, the molecular modeling program identifies target sequences that are most compatible with profiles representing the structural environments of the residues in known threedimensional protein structures. (See, e.g., Eisenberg et al., U.S. Patent No. 5,436,850 issued July 25, 20 1995). In another technique, the known three-dimensional structures of proteins in a given family are superimposed to define the structurally conserved regions in that family. This protein modeling technique also uses the known three-dimensional structure of a homologous protein to approximate the structure of the polypeptide codes of SEQ ID NOS. 812-1599. (See e.g., Srinivasan, et al., U.S. Patent No. 5,557,535 issued September 17, 1996). Conventional homology modeling techniques 25 have been used routinely to build models of proteases and antibodies. (Sowdhamini et al., Protein Engineering 10:207, 215 (1997)). Comparative approaches can also be used to develop threedimensional protein models when the protein of interest has poor sequence identity to template proteins. In some cases, proteins fold into similar three-dimensional structures despite having very weak sequence identities. For example, the three-dimensional structures of a number of helical 30 cytokines fold in similar three-dimensional topology in spite of weak sequence homology.

The recent development of threading methods now enables the identification of likely folding patterns in a number of situations where the structural relatedness between target and template(s) is not detectable at the sequence level. Hybrid methods, in which fold recognition is performed using Multiple Sequence Threading (MST), structural equivalencies are deduced from the threading output—using a distance geometry program DRAGON to construct a low resolution model, and a full-atom representation is constructed using a molecular modeling package such as QUANTA.

According to this 3-step approach, candidate templates are first identified by using the novel fold recognition algorithm MST, which is capable of performing simultaneous threading of multiple aligned sequences onto one or more 3-D structures. In a second step, the structural equivalencies obtained from the MST output are converted into interresidue distance restraints and fed into the distance geometry program DRAGON, together with auxiliary information obtained from secondary structure predictions. The program combines the restraints in an unbiased manner and rapidly generates a large number of low resolution model confirmations. In a third step, these low resolution model confirmations are converted into full-atom models and subjected to energy minimization using the molecular modeling package QUANTA. (See e.g., Aszódi et al., Proteins:Structure, Function, and Genetics, Supplement 1:38-42 (1997)).

The results of the molecular modeling analysis may then be used in rational drug design techniques to identify agents which modulate the activity of the polypeptide codes of SEQ ID NOS. 812-1599.

Accordingly, another aspect of the present invention is a method of identifying a feature

within the nucleic acid codes of SEQ ID NOs. 24-811 and 1600-1622 or the polypeptide codes of

SEQ ID NOS. 812-1599 comprising reading the nucleic acid code(s) or the polypeptide code(s)

through the use of a computer program which identifies features therein and identifying features

within the nucleic acid code(s) or polypeptide code(s) with the computer program. In one

embodiment, computer program comprises a computer program which identifies open reading

frames. In a further embodiment, the computer program identifies structural motifs in a polypeptide

sequence. In another embodiment, the computer program comprises a molecular modeling program.

The method may be performed by reading a single sequence or at least 2, 5, 10, 15, 20, 25, 30, or 50 of
the nucleic acid codes of SEQ ID NOs. 24-811 and 1600-1622 or the polypeptide codes of SEQ ID

NOS. 812-1599 through the use of the computer program and identifying features within the nucleic

acid codes or polypeptide codes with the computer program.

The nucleic acid codes of SEQ ID NOs. 24-811 and 1600-1622 or the polypeptide codes of SEQ ID NOS. 812-1599 may be stored and manipulated in a variety of data processor programs in a variety of formats. For example, the nucleic acid codes of SEQ ID NOs. 24-811 and 1600-1622 or the polypeptide codes of SEQ ID NOS. 812-1599 may be stored as text in a word processing file, such as MicrosoftWORD or WORDPERFECT or as an ASCII file in a variety of database programs familiar to those of skill in the art, such as DB2, SYBASE, or ORACLE. In addition, many computer programs and databases may be used as sequence comparers, identifiers, or sources of reference nucleotide or polypeptide sequences to be compared to the nucleic acid codes of SEQ ID NOs. 24-811 and 1600-1622 or the polypeptide codes of SEQ ID NOS-812-1599. The following list is interded not to limit the invention but to provide guidance to programs and databases which are useful with the nucleic acid codes of SEQ ID NOs. 24-811 and 1600-1622 or the polypeptide codes of SEQ ID NOS. 812-1599. The programs and databases which may be used include, but are not limited to: MacPattern (EMBL),

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DiscoveryBase (Molecular Applications Group), GeneMine (Molecular Applications Group), Look (Molecular Applications Group), MacLook (Molecular Applications Group), BLAST and BLAST2 (NCBI), BLASTN and BLASTX (Altschul et al, J. Mol. Biol. 215: 403 (1990)), FASTA (Pearson and Lipman, Proc. Natl. Acad. Sci. USA, 85: 2444 (1988)), FASTDB (Brutlag et al. Comp. App. Biosci. 5 6:237-245, 1990), Catalyst (Molecular Simulations Inc.), Catalyst/SHAPE (Molecular Simulations Inc.), Cerius².DBAccess (Molecular Simulations Inc.), HypoGen (Molecular Simulations Inc.), Insight II, (Molecular Simulations Inc.), Discover (Molecular Simulations Inc.), CHARMm (Molecular Simulations Inc.), Felix (Molecular Simulations Inc.), DelPhi, (Molecular Simulations Inc.), QuanteMM, (Molecular Simulations Inc.), Homology (Molecular Simulations Inc.), Modeler (Molecular Simulations 10 Inc.), ISIS (Molecular Simulations Inc.), Quanta/Protein Design (Molecular Simulations Inc.), WebLab (Molecular Simulations Inc.), WebLab Diversity Explorer (Molecular Simulations Inc.), Gene Explorer (Molecular Simulations Inc.), SeqFold (Molecular Simulations Inc.), the EMBL/Swissprotein database, the MDL Available Chemicals Directory database, the MDL Drug Data Report data base, the Comprehensive Medicinal Chemistry database, Derwents's World Drug Index database, the 15 BioByteMasterFile database, the Genbank database, and the Genseqn database. Many other programs and data bases would be apparent to one of skill in the art given the present disclosure.

Motifs which may be detected using the above programs include sequences encoding leucine zippers, helix-turn-helix motifs, glycosylation sites, ubiquitination sites, alpha helices, and beta sheets, signal sequences encoding signal peptides which direct the secretion of the encoded proteins, 20 sequences implicated in transcription regulation such as homeoboxes, acidic stretches, enzymatic active sites, substrate binding sites, and enzymatic cleavage sites.

EXAMPLE 62

Methods of Making Nucleic Acids

The present invention also comprises methods of making the EST-related nucleic acids, fragments of EST-related nucleic acids, positional segments of the EST-related nucleic acids, or fragments of positional segments of the EST-related nucleic acids. The methods comprise sequentially linking together nucleotides to produce the nucleic acids having the preceding sequences. A variety of methods of synthesizing nucleic acids are known to those skilled in the art.

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In many of these methods, synthesis is conducted on a solid support. These included the 3' phosphoramidite methods in which the 3' terminal base of the desired oligonucleotide is immobilized on an insoluble carrier. The nucleotide base to be added is blocked at the 5' hydroxyl and activated at the 3' hydroxyl so as to cause coupling with the immobilized nucleotide base. Debiceking of the new immobilized nucleotide compound and repetition of the cycle will produce the desired 35 polynucleotide. Alternatively, polynucleotides may be prepared as described in U.S. Patent No. 5,049,656. In some embodiments, several polynucleotides prepared as described above are ligated

together to generate longer polynucleotides having a desired sequence.

EXAMPLE 63

Methods of Making Polypeptides

The present invention also comprises methods of making the polynucleotides encoded by EST-5 related nucleic acids, fragments of EST-related nucleic acids, positional segments of the EST-related nucleic acids, or fragments of positional segments of the EST-related nucleic acids and methods of making the EST-related polypeptides, fragments of EST-related polypeptides, positional segments of EST-related polypeptides, or fragments of EST-related polypeptides. The methods comprise sequentially linking together amino acids to produce the nucleic polypeptides having the preceding 10 sequences. In some embodiments, the polypeptides made by these methods are 150 amino acid or less in length. In other embodiments, the polypeptides made by these methods are 120 amino acids or less in length.

A variety of methods of making polypeptides are known to those skilled in the art, including methods in which the carboxyl terminal amino acid is bound to polyvinyl benzene or another suitable 15 resin. The amino acid to be added possesses blocking groups on its amino moiety and any side chain reactive groups so that only its carboxyl moiety can react. The carboxyl group is activated with carbodiimide or another activating agent and allowed to couple to the immobilized amino acid. After removal of the blocking group, the cycle is repeated to generate a polypeptide having the desired sequence. Alternatively, the methods described in U.S. Patent No. 5,049,656 may be used.

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As discussed above, the EST-related nucleic acids, fragments of the EST-related nucleic acids, positional segments of the EST-related nucleic acids, or fragments of positional segments of the EST-related nucleic acids can be used for various purposes. The polynucleotides can be used to express recombinant protein for analysis, characterization or therapeutic use; production of secreted polypeptides or chimeric polypeptides, antibody production, as markers for tissues in which the 25 corresponding protein is preferentially expressed (either constitutively or at a particular stage of tissue differentiation or development or in disease states); as molecular weight markers on Southern gels; as chromosome markers or tags (when labeled) to identify chromosomes or to map related gene positions; to compare with endogenous DNA sequences in patients to identify potential genetic disorders; as probes to hybridize and thus discover novel, related DNA sequences; as a source of information to derive PCR 30 primers for genetic fingerprinting; for selecting and making oligomers for attachment to a "gene chip" or other support, including for examination for expression patterns; to raise anti-protein antibodies using DNA immunization techniques; and as an antigen to raise anti-DNA antibodies or elicit another immune response. Where the polynucleotide encodes a protein or polypeptide which binds or potentially binds to another protein or polypeotide (such as, for example, in a receptor-ligand interaction), the polynucleotide 35 can also be used in interaction trap assays (such as, for example, that described in Gyuris et al., Cell 75:791-803 (1993)) to identify polynucleotides encoding the other protein or polypeptide with which

binding occurs or to identify inhibitors of the binding interaction.

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The proteins or polypeptides provided by the present invention can similarly be used in assays to determine biological activity, including in a panel of multiple proteins for high-throughput screening; to raise antibodies or to elicit another immune response; as a reagent (including the labeled reagent) in assays designed to quantitatively determine levels of the protein (or its receptor) in biological fluids; as markers for tissues in which the corresponding protein is preferentially expressed (either constitutively or at a particular stage of tissue differentiation or development or in a disease state); and, of course, to isolate correlative receptors or ligands. Where the protein or polypeptide binds or potentially binds to another protein or polypeptide (such as, for example, in a receptor-ligand interaction), the protein can be used to identify the other protein with which binding occurs or to identify inhibitors of the binding interaction. Proteins or polypeptides involved in these binding interactions can also be used to screen for peptide or small molecule inhibitors or agonists of the binding interaction.

Any or all of these research utilities are capable of being developed into reagent grade or kit format for commercialization as research products.

Methods for performing the uses listed above are well known to those skilled in the art.

References disclosing such methods include without limitation "Molecular Cloning; A Laboratory Manual," 2d ed., Cold Spring Harbor Laboratory Press, Sambrook, J., E.F. Fritsch and T. Maniatis eds., 1989, and "Methods in Enzymology; Guide to Molecular Cloning Techniques," Academic Press, Berger, S.L. and A.R. Kimmel eds., 1987.

Polynucleotides and proteins or polypeptides of the present invention can also be used as

nutritional sources or supplements. Such uses include without limitation use as a protein or amino acid
supplement, use as a carbon source, use as a nitrogen source and use as a source of carbohydrate. In
such cases the protein or polynucleotide of the invention can be added to the feed of a particular
organism or can be administered as a separate solid or liquid preparation, such as in the form of powder,
pills, solutions, suspensions or capsules. In the case of microorganisms, the protein or polynucleotide of
the invention can be added to the medium in or on which the microorganism is cultured.

Although this invention has been described in terms of certain preferred embodiments, other embodiments which will be apparent to those of ordinary skill in the art in view of the disclosure herein are also within the scope of this invention. Accordingly, the scope of the invention is intended to be limited only by reference to the appended claims.

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Sequence Listing Free Text

The following free text appears in the accompanying Sequence Listing:

Von Heijne matrix

-score

35 sequence

name

martinspector prediction

CLAIMS

- A purified nucleic acid comprising a sequence selected from the group consisting of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622 and sequences complementary to the sequences of
 SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622.
 - 2. A purified nucleic acid comprising at least 15 consecutive nucleotides of a sequence selected from the group consisting of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622 and sequences complementary to the sequences of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622.
 - 3. A purified or isolated polypeptide comprising a sequence selected from the group consisting of the sequences of SEQ ID NOs. 812-1599.
 - 4. A method of making a cDNA comprising the steps of:

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- a) contacting a collection of mRNA molecules from human cells with a primer comprising at least 15 consecutive nucleotides of a sequence selected from the group consisting of the sequences complementary to SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622;
 - b) hybridizing said primer to an mRNA in said collection that encodes said protein;
- c) reverse transcribing said hybridized primer to make a first cDNA strand from said mRNA;
 - d) making a second cDNA strand complementary to said first cDNA strand; and
 - e) isolating the resulting cDNA comprising said first cDNA strand and said second cDNA strand.
 - 5. A method of making a cDNA comprising the steps of:
 - a) obtaining a cDNA comprising a sequence selected from the group consisting of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622;
- b) contacting said cDNA with a detectable probe comprising at least 15 consecutive nucleotides of a sequence selected from the group consisting of SEQ ID NOs. 24-811 and SEQ ID
 NOs. 1600-1622 and the sequences complementary to SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622 under conditions which permit said probe to hybridize to said cDNA;
 - c) identifying a cDNA which hybridizes to said detectable probe; and
 - d) isolating said cDNA which hybridizes to said probe.
 - 6. A method of making a cDNA comprising the steps of:
 - a) contacting a collection of mRNA molecules from human cells with a first primer capable of hybridizing to the polyA tail of said mRNA;
 - b) hybridizing said first primer to said polyA tail;

- c) reverse transcribing said mRNA to make a first cDNA strand;
- d) making a second cDNA strand complementary to said first cDNA strand using at least one primer comprising at least 15 consecutive nucleotides of a sequence selected from the group consisting of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622; and
- e) isolating the resulting cDNA comprising said first cDNA strand and said second cDNA strand.
 - 7. A method of making a polypeptide comprising the steps of:
- a) obtaining a cDNA which encodes a polypeptide encoded by a nucleic acid comprising
 a sequence selected from the group consisting of SEQ ID NOs. 24-811 or a cDNA which encodes a polypeptide comprising at least 10 consecutive amino acids of a polypeptide encoded by a sequence selected from the group consisting of SEQ ID NOs. 24-811;
 - b) inserting said cDNA in an expression vector such that said cDNA is operably linked to a promoter;
 - c) introducing said expression vector into a host cell whereby said host cell produces the protein encoded by said cDNA; and
 - d) isolating said protein.

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- 8. In an array of discrete ESTs or fragments thereof of at least 15 nucleotides in length, the improvement comprising inclusion in said array of at least one sequence selected from the group consisting of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622, the sequences complementary to the sequences of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622 and fragments comprising at least 15 consecutive nucleotides of said sequence.
- 9. The array of Claim 8 including therein at least five sequences selected from the group consisting of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622, the sequences complementary to the sequences of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622 and fragments comprising at least 15 consecutive nucleotides of said sequences.
- 10. An enriched population of recombinant nucleic acids, said recombinant nucleic acids comprising an insert nucleic acid and a backbone nucleic acid, wherein at least 5% of said insert nucleic acids in said population comprise a sequence selected from the group consisting of SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622, the sequences complementary to SEQ ID NOs. 24-811 and SEQ ID NOs. 1600-1622 and fragments comprising at least 15 consecutive nucleotides of said sequences.
 - 11. An antibody composition capable of selectively binding to an epitope-containing fragment of a polypeptide comprising a contiguous span of at least 8 amino acids of any of SEQ ID NOs. 812-1599, wherein said antibody is polyclonal or monoclonal.

- 12. A computer readable medium having stored thereon a sequence selected from the group consisting of a nucleic acid code of SEQ ID NOs. 24-811 and 1600-1622 and a polypeptide code of SEQ ID NOs. 812-1599.
- 13. A computer system comprising a processor and a data storage device wherein said data storage device has stored thereon a sequence selected from the group consisting of a nucleic acid code of SEQID NOs. 24-811 and 1600-1622 and a polypeptide code of SEQ ID NOs. 812-1599.
- 14. The computer system of Claim 13 further comprising a sequence comparer and a data storage device having reference sequences stored thereon.
 - 15. The computer system of Claim 14 wherein said sequence comparer comprises a computer program which indicates polymorphisms.
 - 16. The computer system of Claim 13 further comprising an identifier which identifies features in said sequence.
- 17. A method for comparing a first sequence to a reference sequence wherein said first
 20 sequence is selected from the group consisting of a nucleic acid code of SEQID NOs. 24-811 and
 1600-1622 and a polypeptide code of SEQ ID NOs. 812-1599 comprising the steps of:
 - a) reading said first sequence and said reference sequence through use of a computer program which compares sequences; and
- b) determining differences between said first sequence and said reference sequence with 25 said computer program.
 - 18. The method of Claim 17, wherein said step of determining differences between the first sequence and the reference sequence comprises identifying polymorphisms.
- 19. A method for identifying a feature in a sequence selected from the group consisting of a nucleic acid code of SEQID NOs. 24-811 and 1600-1622 and a polypeptide code of SEQ ID NOs. 812-1599 comprising the steps of:
 - a) reading said sequence through the use of a computer program which identifies features in sequences; and
 - b) identifying features in said sequence with said computer program.
 - 20. A vector comprising a nucleic acid according to either Claims 1 or 2.
 - 21. A host cell containing a nucleic acid of Claim 20.

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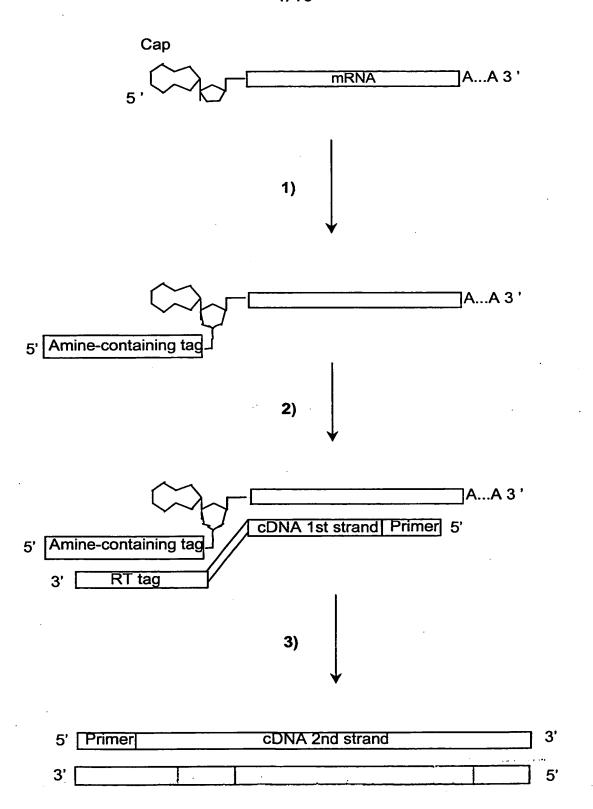


Figure 1

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Minimum signal peptide score	false positive rate	false negative rate	proba(0.1)	proba(0.2)
3,5	0,121	0,036	0,467	0,664
4	0,096	0,06	0,519	0,708
4,5	0,078	0,079	0,565	0,745
5	0,062	0,098	0,615	0,782
5,5	0,05	0,127	0,659	0,813
6	0,04	0,163	0,694	0,836
6,5	0,033	0,202	0,725	0,855
7	0,025	0,248	0,763	0,878
7,5	0,021	0,304	0,78	0,889
8	0,015	0,368	0,816	0,909
8,5	0,012	0,418	0,836	0,92
9	0,009	0,512	0,856	0,93
9,5	0,007	0,581	0,863	0,934
10	0,006	0,679	0,835	0,919

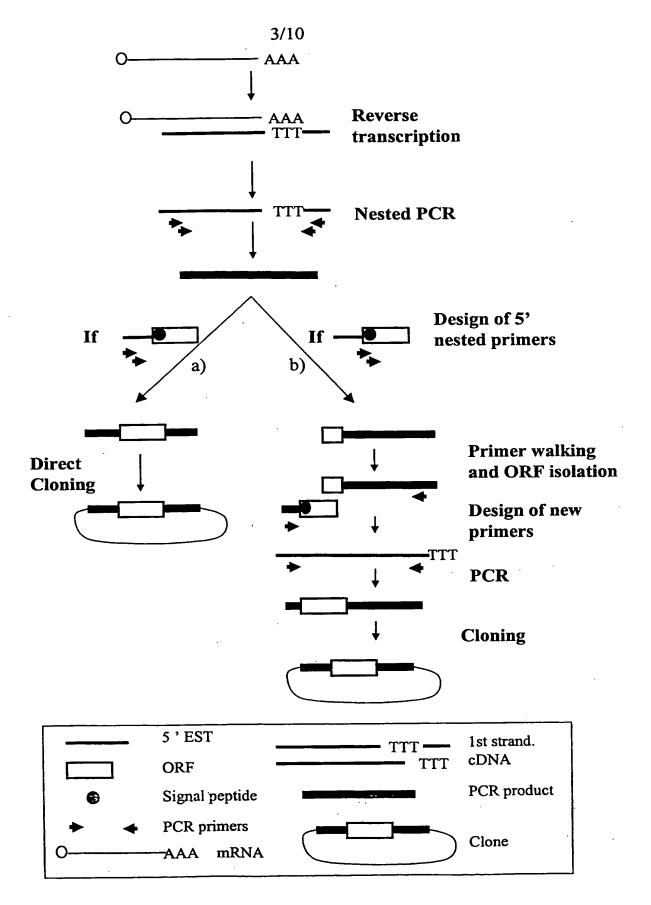


Figure 3

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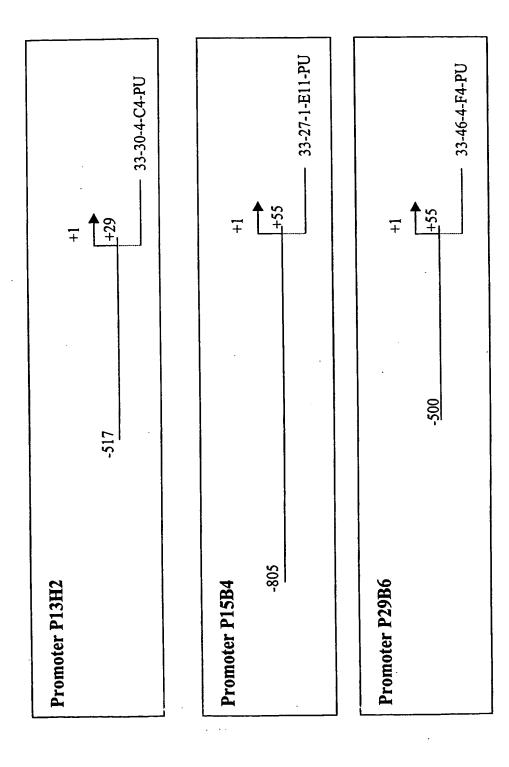


Figure 4

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Promoter sequence P13H2 (546 bp):

		Orient			
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CMYB_01	-502	+	0.983	9	TGTCAGTTG
MYOD_Q6	-501	-	0.961	10	CCCAACTGAC
S8_01	-444		0.960	. 11	AATAGAATTAG
S8_01	-425		0.966	11	AACTAAATTAG
DELTAEF1_01	-390		0.960	11	GCACACCTCAG
	-364		0.964		AGATAAATCCA
GATA_C	-349		0.958		CTTCAGTTG
CMYB_01	-343		0.959		TTGTAGATAGGACA
GATA1_02	-339		0.953		AGATAGGACAT
GATA_C	-235		0.973		CATAACAGATGGTAAG
TAL1ALPHAE47_01	-235		0.983	16	CATAACAGATGGTAAG
TAL1BETAE47_01			0.978	16	CATAACAGATGGTAAG
TAL1BETAITF2_01	-235		0.954	10	ACCATCTGTT
MYOD_Q6	-232		0.953		TCAAGATAAAGTA
GATA1_04	-217				AGTTGGGAATTCC
IK1_01	-126		0.963		AGTTGGGAATTC
IK2_01	-126		0.985		TGGGAATTCC
CREL_01	-123		0.962		TCAGTGATATGGCA
GATA1_02	-96		0.950		
SRY_02	-41		0.951	12	TAAAACAAAACA
E2F_02	-33		0.957		TITAGCGC
MZF1_01	-5	-	0.975	8	TGAGGGGA

Promoter sequence P15B4 (861bp):

_	· • -	4
u	rıe	nt

Matrix	Position	ation	Score	Length	Sequence
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	-684	+	0.994	9	TGACCGTTG
CMYB_01	-682	-	0.985	9	TCCAACGGT
VMYB_02	-673	+	0.968	9	TTCCTGGAA
STAT_01	-673	-	0.951	9	TTCCAGGAA
STAT_01	-556		0.956	8	TTGGGGGA
MZF1_01	-350 -451	· +	0.965	12	GAATGGGATTTC
IK2_01			0.986	8	AGAGGGGA
MZF1_01	-424	+		12	GAAAACAAAACA
SRY_02	-398		0.955		GAAGGGGA
MZF1_01	-216	+	0.960	8	
MYOD_Q6	-190	+	0.981	10	AGCATCTGCC
DELTAEF1_01	-176	+	0.958	11	TCCCACCTTCC
S8_01	5	-	0.992	11	GAGGCAATTAT
MZF1 01	16	-	0.986	8	AGAGGGGA

Promoter sequence P29B6 (555 bp):

Oriont

		Ottent			
Matrix	Position	ation	Score	Length	Sequence
ARNT_01	-311	+	0.964	16	GGACTCACGTGCTGCT
NMYC_01	-309	+	0.965	12	ACTCACGTGCTG
USF_01	-309	+	0.985	12	ACTCACGTGCTG
USF_01	-309	_	0.985	12	CAGCACGTGAGT
NMYC_01	-309		0.956	12	CAGCACGTGAGT
	-309	_	0.972	12	CAGCACGTGAGT
MYCMAX_02	-307		0.997	8	1CACGTGC
USF_C	-307	-	0.991	8	GCACGTGA
USF_C	-292		0.968	8	CATGGGGA
MZF1_01	-105		0.963	14	CTCTCCGGAAGCCT
ELK1_02	-102		0.974	10	TCCGGAAGCC
CETS1P54_01	-42		0.963	11	AGTGACTGAAC
AP1_Q4 AP1FJ_Q2	-42		0.961	11	AGTGACTGAAC
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Figure \$

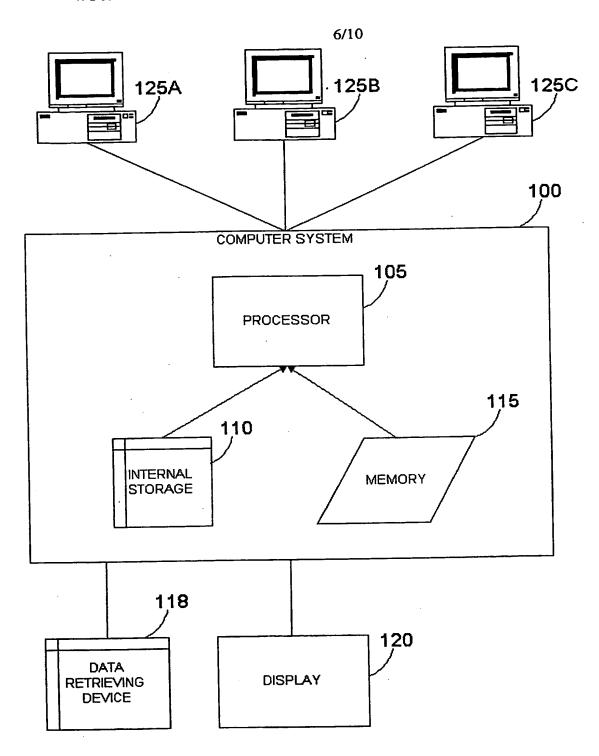


FIGURE 6

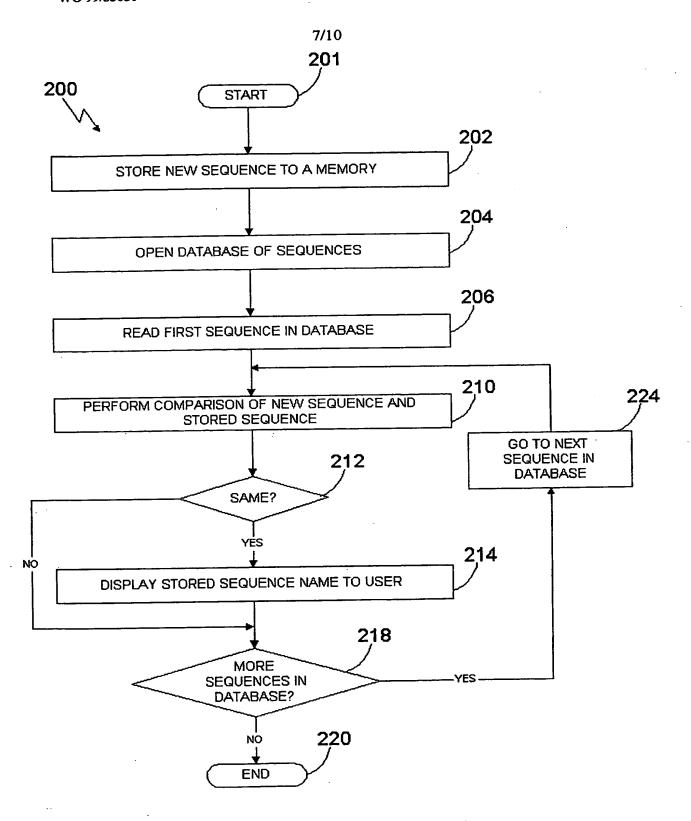
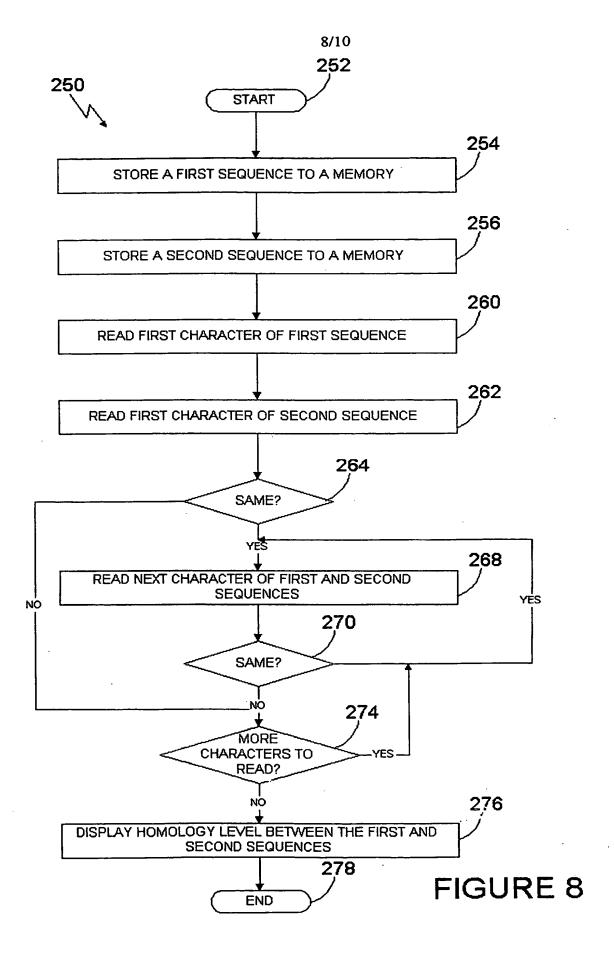
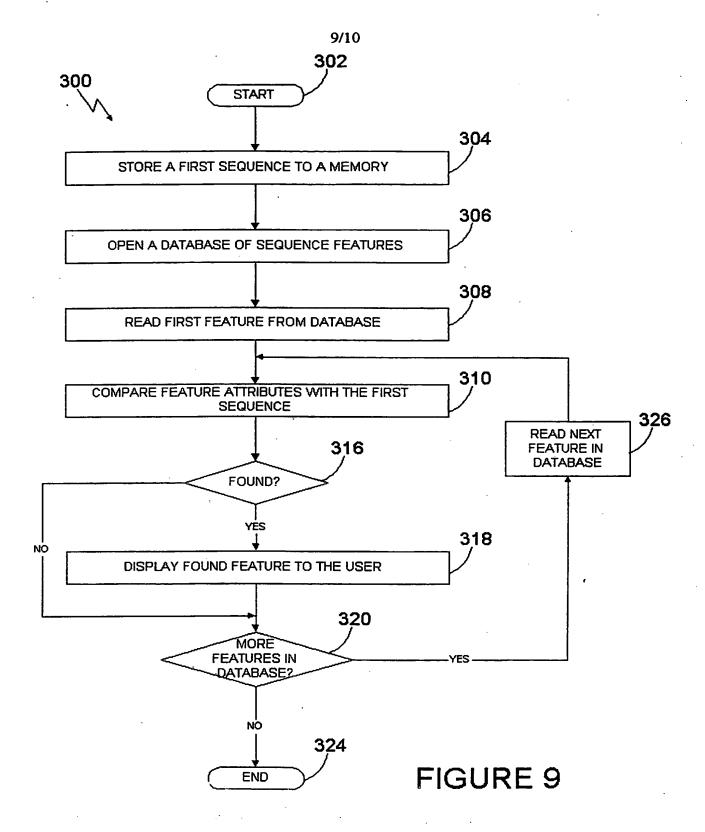


FIGURE 7





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* tRNA	FASTA	both	•	08	09	
rRNA	BLASTN	both	S=108	08	40	
mtRNA	BLASTN	both	S=108	80	40	
Procaryotic	BLASTN	both	S=144	8	40	
Fungal	BLASTN	both	S=144	06	40	
§ Alu	BLASTN	both	S=72, B=5	70	40	max 5 matches, masking
, L1	BLASTN	both	S=72, B=5	20	40	max 5 matches, masking
Repeats	BLASTN	both	S=72	70	40	masking
			W=6, S=10, E=1000,	-		
PolyA	BLAST2N	top	N=12	. 06	10	in the last 100 nucleotides
Folyadenylation signal	•	top	AATAAA allowing 1 mismatch	msim 1 mism	natch	in the 50 nucleotides before the 5' end of the polA
Vertebrate	BLASTN then FASTA	both	•	90 then 70	30	first BLASTN, then FASTA on maching sequences
ESTs	BLAST2N	both	•	96	30	
Genesed	BLASTN	both	W=8, B=10	96	30	
ORF	BLASTP	top	W=8, B=10	•	•	on ORF proteins, max 10 matches
Proteins	BLASTX	top	E = 0.001	70	30	

Figure 10

SEQUENCE LISTING

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Pro Ile Val Lys Phe Pro Gly Ala Val Asp Gly Ala Thr Tyr Ile Leu . 60 65 gtg atg gtg gat cca gat gcc cct agc aga gca gaa ccc aga cag aga 338 Val Met Val Asp Pro Asp Ala Pro Ser Arg Ala Glu Pro Arg Gln Arg 80 ttc tgg aga cat tgg ctg gta aca gat atc aag ggc gcc gac ctg aag 386 Phe Trp Arg His Trp Leu Val Thr Asp Ile Lys Gly Ala Asp Leu Lys aaa ggg aag att cag ggc cag gag tta tca gcc tac cag gct ccc tcc 434 Lys Gly Lys Ile Gln Gly Gln Glu Leu Ser Ala Tyr Gln Ala Pro Ser 110 cca ccg gca cac agt ggc ttc cat cgc tac cag ttc ttt gtc tat ctt 482 Pro Pro Ala His Ser Gly Phe His Arg Tyr Gln Phe Phe Val Tyr Leu 125 cag gaa gga aag gtc atc tct ctc ctt ccc aag gaa aac aaa act cga -530 Gln Glu Gly Lys Val Ile Ser Leu Leu Pro Lys Glu Asn Lys Thr Arg 145 140 ggc tct tgg aaa atg gac aga ttt ctg aac cgt ttc cac ctg ggc gaa 578 Gly Ser Trp Lys Met Asp Arg Phe Leu Asn Arg Phe His Leu Gly Glu 155 160 cct gaa gca agc acc cag ttc atg acc cag aac tac cag gac tca cca 626 Pro Glu Ala Ser Thr Gln Phe Met Thr Gln Asn Tyr Gln Asp Ser Pro 170 175 acc ctc cag gct ccc aga gaa agg gcc agc gag ccc aag cac aaa aac 674 Thr Leu Gln Ala Pro Arg Glu Arg Ala Ser Glu Pro Lys His Lys Asn 190 195 cag gcg gag ata gct gcc tgc tagatagccg gctttgccat ccgggcatgt 725 Gln Ala Glu Ile Ala Ala Cys 205 ggccacactg cccaccaccg acgatgtggg tatggaaccc cctctggata cagaacccct 785 826 <210> 8 <211> 227 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -22..-1 <223> score 8.5 seq AALLLGLMMVVTG/DE <400> 8 Met Gly Trp Thr Met Arg Leu Val Thr Ala Ala Leu Leu Gly Leu Met Met Val Val Thr Gly Asp Glu Asp Glu Asn Ser Pro Cys Ala His Glu Ala Leu Leu Asp Glu Asp Thr Leu Phe Cys Gln Gly Leu Glu Val 20 Phe Tyr Pro Glu Leu Gly Asn Ile Gly Cys Lys Val Val Pro Asp Cys 35 Asn Asn Tyr Arg Gln Lys Ile Thr Ser Trp Met Glu Pro Ile Val Lys 50 Phe Pro Gly Ala Val Asp Gly Ala Thr Tyr Ile Leu Val Met Val Asp 65... 70 Pro Asp Ala Pro Ser Arg Ala Glu Pro Arg Gln Arg Phe Trp Arg His 80 85 Trp Leu Val Thr Asp Ile Lys Gly Ala Asp Leu Lys Lys Gly Lys Ile

100

Gln Gly Gln Glu Leu Ser Ala Tyr Gln Ala Pro Ser Pro Pro Ala His 110 115 120 Ser Gly Phe His Arg Tyr Gln Phe Phe Val Tyr Leu Gln Glu Gly Lys

6 125 130 Val Ile Ser Leu Leu Pro Lys Glu Asn Lys Thr Arg Gly Ser Trp Lys 145 150 Met Asp Arg Phe Leu Asn Arg Phe His Leu Gly Glu Pro Glu Ala Ser 165 160 Thr Gln Phe Met Thr Gln Asn Tyr Gln Asp Ser Pro Thr Leu Gln Ala 175 180 Pro Arg Glu Arg Ala Ser Glu Pro Lys His Lys Asn Gln Ala Glu Ile 195 Ala Ala Cys 205 <210> 9 <211> 852 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 229..735 <221> sig_peptide <222> 229..492 <223> score 6.7 seq VFALSSFLNKASA/VY <400> 9 aatgactggc agtggcatca gcgatggcgg ctgcgtcggg gtcggttctg cagcgctgta 60 tegtgtegee ggeagggagg catagegeet etetgatett cetgeatgge teaggtgatt 120 ctggacaagg attaagaatg tggatcaagc aggtttttaa atcaagattt aacattccaa 180 cacataaaaa ttatttatcc aacagctcct cccagatcat atactcct atg aaa gga 237 Met Lys Gly gga atc tcc aat gta tgg ttt gac aga ttt aaa ata acc aat gac tgc 285 Gly Ile Ser Asn Val Trp Phe Asp Arg Phe Lys Ile Thr Asn Asp Cys -80 -75 cca gaa cac ctt gaa tca att gat gtc atg tgt caa gtg ctt act gat 333 Pro Glu His Leu Glu Ser Ile Asp Val Met Cys Gln Val Leu Thr Asp -65 -60 ttg att gat gaa gaa gta aaa agt ggc atc aag aag aac agg ata tta 381 Leu Ile Asp Glu Glu Val Lys Ser Gly Ile Lys Lys Asn Arg Ile Leu -50 -45 ata gga gga ttc tct atg gga gga tgc atg gca atg cat tta gca tat 429 Ile Gly Gly Phe Ser Met Gly Gly Cys Met Ala Met His Leu Ala Tyr -30 -25 aga aat cat caa gat gtg gca gga gta ttt gct ctt tct agt ttt ctg 477 Arg Asn His Gln Asp Val Ala Gly Val Phe Ala Leu Ser Ser Phe Leu -15 aat aaa gca tct gct gtt tac cag gct ctt cag aag agt aat ggt gta 525 Asn Lys Ala Ser Ala Val Tyr Gln Ala Leu Gln Lys Ser Asn Gly Val ctt cct gaa tta ttt cag tgt cat ggt act gca gat gag tta gtt ctt 573 Leu Pro Glu Leu Phe Gln Cys His Gly Thr Ala Asp Glu Leu Val Leu 15 20 cat tot tgg gca gaa gag aca aac toa atg tta aaa tot ota gga gtg 621 His Ser Trp Ala Glu Glu Thr Asn Ser Met Leu Lys Ser Leu Gly Val 35 acc acg aag ttt cat agt ttt cca aat gtt tac cat gag cta agc aaa 669 Thr Thr Lys Phe His Ser Phe Pro Asn Val Tyr His Glu Leu Ser Lys. 50 act gag tta gac ata ttg aag tta tgg att ctt aca aag ctg cca gga 717 Thr Glu Leu Asp Ile Leu Lys Leu Trp Ile Leu Thr Lys Leu Pro Gly 70 gaa atg gaa aaa caa aaa tgaatgaatc aagagtgatt tgttaatgta 765

149

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Arg Lys Cys Ser Val Phe His Leu Phe Val Ala Cys Leu Ser Leu Gly

PCT/IB99/00712 WO 99/53051 8

			-20					-15	Ū				-10			
ttc	ttc	tcc		ctc	taa	cta	caq		agc	tac	tct	aaa		ata	acc	197
									Ser							
~~~		_		~~~	<b>C</b> 3 3	~~~	_	<i>~~~</i>	acc	toa	2	cct	~~~	cat	acc	245
									Thr							243
10	Ата	vai	Arg	GIY	15	GIY	GIII	GIU	1111	20	GIY	PIO	PIO	Arg	25	
	~~~	<b>~~</b>	a2a	cca		cct	asa	cac	taa		<b>~</b> 22	asc.		tee		293
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Cys	PIO	PIO	GIU	30	110	110	Giu	1113	35	Olu	Giu	АЗР	ALG	40	112	
aac	ccc	cac	cac		aca	ata	cta	ata	ccc	ttc	cac	паа	cac		aaa	341
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Gry	PIO	птэ	45	пец	AIG	val	Бец	50	PIO	FIIC	Arg	Giu	55	FIIC	GIU	
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		_	_					_	Arg			_	_		-	303
GIU	пеп	60	vai	FIIC	Vai	FIO	65	MEC	Arg	AT 9	FIIC	70	Ser	Arg	Буб	
220	atc		cac	cac.	atc	tac		ctc	aac	cad	ata		cac	ttc	agg	437
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_	_	_			_	_		_	Val	_	_					
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			125		-			130					135			•
tcc	cca	gag		cac	cct	ctc	tac		tac	aaq	acc	tat	qtc	qqc	ggc	629
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Asp	Gln	Lys	Arg	Ile	Ala	Ala		Lys	Gln	Glu	Gln	Phe	Lys	Val	Asp	
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Arg		Gly	Gly	Leu	Asn		Val	Lys	Tyr	His		Ala	Ser	Arg	Thr	
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	Leu	Ser	Val	Gly		Ala	Pro	Cys	Thr		Leu	Asn	Ile	Met		
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Asp	Cys	Asp	Lys		Ala	Tha	Pro	Trp	Cys	Thr	Phe	Ser				
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Val Ala Pro Val	ccc tgt aca Pro Cys Thr	acc ctg ctg c Thr Leu Leu P -130	-140 cc tgt caa acc ctg ttc ro Cys Gln Thr Leu Phe -125	
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11

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22

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Leu Pro Pro Pro Gly Ser Cys Ala Gly Arg Arg Ser Pro Xaa Thr Pro

gac gag tot acc cca cct ccc cgg aag aag aag gat att cgc gat 195 Asp Glu Ser Thr Pro Pro Pro Arg Lys Lys Lys Asp Ile Arg Asp 10 15 tac aat gat gca gac atg gcg cgt ctt ctg gag caa ggg gag ggg 240 Tyr Asn Asp Ala Asp Met Ala Arg Leu Leu Glu Gln Gly Glu Gly 30 35 <210> 30

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tct gga ttc aat ttt agc act tat gag atg cat tgg atc cgc cag gct Ser Gly Phe Asn Phe Ser Thr Tyr Glu Met His Trp Ile Arg Gln Ala 30

cca ggg aag ggg ccg gag tgg gtn nca tat gtc agt ggt gga ggt gga Pro Gly Lys Gly Pro Glu Trp Val Xaa Tyr Val Ser Gly Gly Gly Gly 50

acc agh nnn aac gev sac tet gtg aag gge ega tte ace ate tee aga-352 Thr Xaa Xaa Asn Ala Xaa Ser Val Lys Gly Arg Phe Thr Ile Ser Arg 65

304

gac aat gcc aac agt ttt gtg tat cta caa atg gac agt ctg cga gtc 400 Asp Asn Ala Asn Ser Phe Val Tyr Leu Gln Met Asp Ser Leu Arg Val

80 gag gac acc gct ctc tat tac tgt gcg aga rgg gat tac gac ttc tgg 448 Glu Asp Thr Ala Leu Tyr Tyr Cys Ala Arg Xaa Asp Tyr Asp Phe Trp 95

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Ser Gly Tyr Tyr

105

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54

445

WO 99/53051 24 <221> CDS <222> 28..111 <221> sig_peptide <222> 28..84 <223> Von Heijne matrix score 13.3999996185303 seq LLLLLSHCTGSLS/QP <400> 31 aactgtgcat gtcaggctgt gtccacc atg gcc tgg act cct ctt ctc ttg Met Ala Trp Thr Pro Leu Leu Leu ctc ctc tct cac tgc aca ggt tcc ctc tcc cag cct gtg ctg act cag 102 Leu Leu Ser His Cys Thr Gly Ser Leu Ser Gln Pro Val Leu Thr Gln

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Glu Asp Thr Ala Val Tyr Tyr Cys Ala Lys Asp Arg Thr Gly Gly

90 95 100 <210> 33

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70

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WO 99/53051 28 <220> <221> CDS <222> 9..161 <221> sig peptide <222> 9..83 <223> Von Heijne matrix score 11.8999996185303 seq WLLLLPLLLGLNA/GA 50 aacttqtc atg gag ctg gca ctg cgg cgc tct ccc gtc ccg cgg tgg ttg Met Glu Leu Ala Leu Arg Arg Ser Pro Val Pro Arg Trp Leu -20 ctq ctg ctg ctg ctg ctg ggc ctg aac gca gga gct gtc att gac 98 Leu Leu Pro Leu Leu Gly Leu Asn Ala Gly Ala Val Ile Asp -5 tgg ccc aca gag gag ggc aag gaa gta tgg gat tat gtg acg gtc cgc 146 Trp Pro Thr Glu Glu Gly Lys Glu Val Trp Asp Tyr Val Thr Val Arg 10 163 aag gat gcc tac atg gg Lys Asp Ala Tyr Met 25 <210> 39 <211> 427 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 35..427 <221> sig_peptide <222> 35..91 <223> Von Heijne matrix score 11.8999996185303 seq FLFLLTCCPGSNS/QA <221> misc feature <222> 138..139 <223> n=a, g, c or t <400> 39 55 tetggeacca ggggteeett ceaatateag eace atg gee tgg act eet ete ttt Met Ala Trp Thr Pro Leu Phe ctg ttc ctc ctc act tgc tgc cca ggg tcc aat tcc cag gct gtg gkg 103 Leu Phe Leu Leu Thr Cys Cys Pro Gly Ser Asn Ser Gln Ala Val Xaa act cag gag ccc ctc act gac tgt gtc ccc cgg ann aca gtc act ctc 151 Thr Gln Glu Pro Leu Thr Asp Cys Val Pro Arg Xaa Thr Val Thr Leu 10 15 1.95 acc tgt ggc too agt att gga gct gtc acc aat ggt cat ttt ccc tac Thr Cys Gly Ser Ser Ile Gly Ala Val Thr Asn Gly His Phe Pro Tyr 25 30

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29 Leu Gly Gly Lys Ala Val Leu Thr Leu Ser Asp Ala Gln Pro Asp Asp 70 gag gct gaa tat tat tgt gtc ctc tcc tat agt ggt ggt cgg ccg gtg 391 Glu Ala Glu Tyr Tyr Cys Val Leu Ser Tyr Ser Gly Gly Arg Pro Val 427 ttc ggc gga ggg acc aag ctg acc gtc cta agt cag Phe Gly Gly Thr Lys Leu Thr Val Leu Ser Gln <210> 40 <211> 97 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 22..96 <221> sig peptide <222> 22..84 <223> Von Heijne matrix score 11.8999996185303 seq LALCLLLGPLAGA/KP agatcaggaa gcaccgggaa g atg cag gcc tgc atg gtg ccg ggg ctg gcc 51 Met Gln Ala Cys Met Val Pro Gly Leu Ala -20 -15 ctc tgc ctc cta ctg ggg cct ctt gca ggg gcc aag cct gtg cag g 97 Leu Cys Leu Leu Gly Pro Leu Ala Gly Ala Lys Pro Val Gln -10 <210> 41 <211> 536 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 266..535 <221> sig peptide <222> 266..307 <223> Von Heijne matrix score 15 seq LLPLLLLPMCWA/VE <400> 41 actittgggg teaegtgete atteegtite cetaecteec ceaacettat ecegeceetg 60 qqqqttcqcq qqcatttttc aggaactttc tttccggctt gagaagccgc cactcccaag 120 atgsagcagg aaccgcggct gctggacaag aggggtgcgg tggatactga cctttgctcc 180 qqcctcqtcq tqaaqacaca gcgcatctcc ccgctgtagg cttcctccca cagaacccgt 240 ttegggeete agagegtetg gtgag atg etg ttg eeg etg etg etg eta 292 Met Leu Leu Pro Leu Leu Leu Leu Leu one atg tgc tgg gcc gtg gag gtc aag agg ecc mgg ggn gto toc Ctc 340 Pro Met Cys Trp Ala Val Glu Val Lys Arg Pro Arg Gly Val Ser Leu 1 acc aat cat tac tac gat gag too aag cot tto acc tgc otg gac 388 Thr Asn His His Phe Tyr Asp Glu Ser Lys Pro Phe Thr Cys Leu Asp 20 15 ggt tcg gcc acc atc cca ttt gat cag gtc aac gat gac tat tgc gac 436 Gly Ser Ala Thr Ile Pro Phe Asp Gln Val Asn Asp Asp Tyr Cys Asp

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Cys Lys Asp Gly Ser Asp Glu Pro Gly Thr Ala Ala Cys Pro Asn Gly
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                                            55
age tte cae tge ace aac act gge tat aag eec etg tat ate eec tee
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                                                                       172
                         Met Gly Thr Gln Glu Gly Trp Xaa Leu Leu
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ctc tgc ctg gct cta tct gga gca gca gaa acc aag ccc cac cca gca
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Leu Cys Leu Ala Leu Ser Gly Ala Ala Glu Thr Lys Pro His Pro Ala
                        -5
gag ggg cag tgg cgg gca gtg gdc gtg gtc cta gac ygt ttc ctg gtg
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Glu Gly Gln Trp Arg Ala Val Xaa Val Val Leu Asp Xaa Phe Leu Val
aag gac svt geg cac egt gga get ete gee age agt gag gac agg gea
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Lys Asp Xaa Ala His Arg Gly Ala Leu Ala Ser Ser Glu Asp Arg Ala
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Ile Thr Cys Arg Ala Ser Xaa Ser Ile Gly Ser Ser Leu Tyr Trp Tyr

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Gln Gln Lys Pro His Gln Ser Pro Lys Leu Val Ile Lys Tyr Ala Ser

45

40

30

196

32													
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gag aaa att cca gtg Glu Lys Ile Pro Val -15	Ser Ala Ph			er Tyr									
act ctg gcc aga gat Thr Leu Ala Arg Asp 1													
aag gac tot cga ccc Lys Asp Ser Arg Pro													
gac caa ctc atc tgg Asp Gln Leu Ile Trp 30													
aag aca agc aac aaa Lys Thr Ser Asn Lys 50				lu Cys									
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cag aaa ttg gca gag Gln Lys Leu Ala Glu 80	cag ttt gt Gln Phe Va 85	al Leu Leu Asn	ctg gtt tat ga Leu Val Tyr G	aa aca 514 Lu Thr									
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PCT/IB99/00712 WO 99/53051

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244 tgg agt tgg gtc cgc cag ccc cca ggg aag gga ctg gag tgg att tcg Trp Ser Trp Val Arg Gln Pro Pro Gly Lys Gly Leu Glu Trp Ile Ser 40 45 gaa atc gat cat/ggt gga aac acc aat tac aac ccg tcc ctc aag agt 292 Glu Ile Asp Hiś Gly Gly Asn Thr Asn Tyr Asn Pro Ser Leu Lys Ser 60 cqa qtc kcc att tct tta gac aag tcc aag aat aag ttc tcc ctg agg 340 Arg Val Xaa Ile Ser Leu Asp Lys Ser Lys Asn Lys Phe Ser Leu Arg ctg acc tct gtg acc gcc gcg gac acc gcc atg tat kac tgt gcg aga 388 Leu Thr Ser Val Thr Ala Ala Asp Thr Ala Met Tyr Xaa Cys Ala Arg 85 ggc ggt gcg bnc agc tcg tcc gct ttt gat gtc tgg ggc cta rgg aca 436 Gly Gly Ala Xaa Ser Ser Ser Ala Phe Asp Val Trp Gly Leu Xaa Thr 105 110 atg gtc atc atc tct tca gcc tc 459 Met Val Ile Ile Ser Ser Ala <210> 48 <211> 437 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 20..436 <221> sig peptide <222> 20..76 <223> Von Heijne matrix score 11 seq TLLLLTVPSWVLS/QV gtgaatectg etetecace atg gae ata ett tgt tee aeg ete etg etm etg 52 Met Asp Ile Leu Cys Ser Thr Leu Leu Leu Leu -15 ack gtc ccg tcc tgg gtc tta tcc car gtc acc ttg arg gaa tct ggt 100 Thr Val Pro Ser Trp Val Leu Ser Gln Val Thr Leu Xaa Glu Ser Gly 1 cct gcg ctg gtg aaa gcc aca cag acc ctc aga ctg acc tgc acc ttc 148 Pro Ala Leu Val Lys Ala Thr Gln Thr Leu Arg Leu Thr Cys Thr Phe 15 tet qqq tte tea ete aqe act aat aqa atq eqt qtq aqt tqq ate eqt 196 Ser Gly Phe Ser Leu Ser Thr Asn Arg Met Arg Val Ser Trp Ile Arg cag ccc cca ggg aag gcc ctg gag tgg ctt gca cgg att gat tgg gat 244 Gln Pro Pro Gly Lys Ala Leu Glu Trp Leu Ala Arg Ile Asp Trp Asp 45 gat tat aag agg tac agc aca tct ctg aag acc agg gtc acc atc tcc 292 Asp Tyr Lys Arg Tyr Ser Thr Ser Leu Lys Thr Arg Val Thr Ile Ser 65 dag gad acg too aaa aac dag gtg atd otg aca atg acc aac gtg gad 340 Lys Asp Thr Ser Lys Asn Gln Val Ile Leu Thr Met Thr Asn Val Asp 80 cct gcg gac aca gcc acc tat tac tgt gca cgc ctt tca acg gca gct 388 Pro Ala Asp Thr Ala Thr Tyr Tyr Cys Ala Arg Leu Ser Thr Ala Ala ace cea eag tit tit gae tie tig gige eag giga gite etg gite tee gite t 437 Thr Pro Gln Phe Phe Asp Phe Trp Gly Gln Gly Val Leu Val Ser Val

115

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Phe Leu Leu Val Ala Ala Pro Arg Trp Val Leu Ser Gln Val Gln

-5 ctg cag gag tcg ggc cca aga ctg gtg aag cct tca cag acc ctg tcc

-10

103

37											
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tgg agt tgg atc cgc cag cac cca ggg cgg ggc ctg gag tgg att ggc Trp Ser Trp Ile Arg Gln His Pro Gly Arg Gly Leu Glu Trp Ile Gly 40 45 50	247										
tac atc tat tac aat tgg agc acc tac tac aat ccg tcc ctc agg agt Tyr Ile Tyr Tyr Asn Trp Ser Thr Tyr Tyr Asn Pro Ser Leu Arg Ser 55 60 65	295										
cga gtt acc atg tca atg gac acg tct aag aac cag ttc tcc ctg aac Arg Val Thr Met Ser Met Asp Thr Ser Lys Asn Gln Phe Ser Leu Asn 70 75 80	343										
ctg aac tct gta act gcc gcg gac acg gsc atg tat tac tgt gcs aga Leu Asn Ser Val Thr Ala Ala Asp Thr Xaa Met Tyr Tyr Cys Ala Arg 85 90 95	391										
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gtc ctg tcc cag gtg cag ctg cag gag tcg ggc cca gga ctg gtg aag Val Leu Ser Gln Val Gln Leu Gln Glu Ser Gly Pro Gly Leu Val Lys  1 5 10	154										
cct tca gag acc ctg tcc ctc acc tgc act gtc tct ggt ggc tcc atc Pro Ser Glu Thr Leu Ser Leu Thr Cys Thr Val Ser Gly Gly Ser Ile 15 20 25	202										
agg act ggt tot tac tac tgg act tgg gtt cgc cag ccc ccc ggg aag Arg Thr Gly Ser Tyr Tyr Trp Thr Trp Val Arg Gln Pro Pro Gly Lys 30 35 40 45	250										
ggc ctg gag tgg att ggc tac att tat tat act ggg gac acc tac tac Gly Leu Glu Trp Ile Gly Tyr Ile Tyr Tyr Thr Gly Asp Thr Tyr Tyr 50 55 60	298										
aac ccg tcc ctc aag agt cga att acc atg tcr cta gac acg tny wag Asn Pro Ser Leu Lys Ser Arg Ile Thr Met Ser Leu Asp Thr Xaa Xaa	346										

38													
65 70 <b>7</b> 5													
aac cag ttc kcc ctg agc ctg acc tct gtg act gtc gca gac acg g  Asn Gln Phe Xaa Leu Ser Leu Thr Ser Val Thr Val Ala Asp Thr  80 85 90	}2												
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•	45												
agt ctt gcc aaa ttt gat gcc cct cga Ser Leu Ala Lys Phe Asp Ala Pro Arg 30 35	72												
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	gat ag Asp Se		aat					acc					ctc		296		
Ser A	ega gt Arg Va 70	l Thr	Ile	Ser	Xaa	Asp 75	Thr	Ser	Lys	Xaa	Gln 80	Leu	Ser	Leu	344		
Arg I	ctg ac Leu Th 85	r Ser	Val	Thr	Xaa 90	Ala	Asp	Thr	Ala	Val 95	Tyr	Tyr	Cys	Ala	392		
Arg I	aag to Lys Se	r Ser	Met	His 105											440		
	tac tt Tyr Ph	_		99											457		
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	cag ga Gln Gl 5														148		
Leu 1	acc tg Thr Cy 20	s Ala	Val	Ser	Gly 25	Gly	Ser	Ile	Ile	Ser 30	Ser	Asn	Trp	Trp	196		
Ser 1	tgg gt Irp Va	l Arg	Gln	Thr 40	Pro	Gly	Lys	Gly	Leu 45	Glu	Trp	Ile	Gly	Glu 50	244		
Ile T	tat ga Tyr Gl	u Asp	Gly 55	Ile	Thr	Asn	Tyr	Asn 60	Pro	Ser	Leu	Lys	Ser 65	Arg	292		
Val I	atc at	e Ser 70	Val	Asp	Lys	Āla	Lys 75	Asn	Gln	Phe	Ser	Leu 80	ŗys	Met	340		
Arg S	tct gt Ser Va 85	l Thr	Ala	Ser	Asp	Thr 90	Ala	Val	Tyr					_	388		
Ser S	agc to Ser Se 100								ca						420		

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tcc cag gtc cag ctt gtg cag tct ggg gct gag gtg aag aag cct ggg 153 Ser Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly	}											
1 5 10 15 gcc tca gtg aag gtt tcc tgc aag gct tct gga tac ayc ttc act ary 201 Ala Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Xaa Phe Thr Xaa	L											
20 25 30 tmt gct atn cat tgg gtg cgc cag gcc ccc gga car agr ctt gag tgg 249 Xaa Ala Xaa His Trp Val Arg Gln Ala Pro Gly Gln Arg Leu Glu Trp	<b>)</b>											
35 40 45 atg ggr tgg atc aac gct gcc amt ggt wam aca awa tat tca cag aas 297 Met Gly Trp Ile Asn Ala Ala Xaa Gly Xaa Thr Xaa Tyr Ser Gln Xaa	,											
50 55 60  ttc cag grc aga gtc acc wtt acc agg gac aca tcc gcg agc aca gtc 345 Phe Gln Xaa Arg Val Thr Xaa Thr Arg Asp Thr Ser Ala Ser Thr Val	;											
65 70 75 tec atg gag etg age age etg aga tet gaa gae aeg get gtg tat tte 393	j											
Ser Met Glu Leu Ser Ser Leu Arg Ser Glu Asp Thr Ala Val Tyr Phe 80 95 95 tgt gcg aga gat tgg gaa att gca gta gta cca act gct ata aac tct 441	Ĺ											
Cys Ala Arg Asp Trp Glu Ile Ala Val Val Pro Thr Ala Ile Asn Ser 100 105 110 tac ggg ttc gac cct ggg gcc agg gaa cct 471	L											
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teegstgewg ceatttgtga gageegetgg eggetgagtg aaagteattt tgaaagaetg 180 ateeaaagaa ga atg gag gee aga gtg gag egt get gtg eag aaa agg eaa 231 Met Glu Ala Arg Val Glu Arg Ala Val Gln Lys Arg Gln -25 -20 -15												
gtc tta ttt ctt tgt gta ttt ctg gga atg tct tgg gct ggc gcc gaa 279 Val Leu Phe Leu Cys Val Phe Leu Gly Met Ser Trp Ala Gly Ala Glu -10 -5 1	,											
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		gtg cag tct gg Val Gln Ser Gl 5			
		tcc tgc aag go Ser Cys Lys Al			
		gtg cga cag go Val Arg Gln Al 40	la Pro Gly G		
		gct dcc aat go Ala Xaa Asn Gl 55			
		acc atg acc ac Thr Met Thr Th 70	hr Asp Thr S		
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gag cag tta cgg cag tgg ggc gca sga ctg ttg aag cct tcg gag acc Glu Gln Leu Arg Gln Trp Gly Ala Xaa Leu Leu Lys Pro Ser Glu Thr 5 10 15	146										
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tgg agc tgg atc cgc cag tcc cca ggg aag ggg ctg gag tgg att ggg Trp Ser Trp Ile Arg Gln Ser Pro Gly Lys Gly Leu Glu Trp Ile Gly 35 40 45	242										
gga atc aat cac agc gga agc acc ctc tcc aac ccg tcc ctc aag agt Gly Ile Asn His Ser Gly Ser Thr Leu Ser Asn Pro Ser Leu Lys Ser 50 55 60 65	290										
cgc gtc gac ctc tca gtt gat gcg tcc aag gac cag gtg tcc ctg agg Arg Val Asp Leu Ser Val Asp Ala Ser Lys Asp Gln Val Ser Leu Arg 70 75 80	338										
ctg aaa ctt gtg acc gcc gcg gac acg gct gtg tac ttc tgc gcg aga Leu Lys Leu Val Thr Ala Ala Asp Thr Ala Val Tyr Phe Cys Ala Arg 85 90 95	386										
ccc cat tac gat atg tcg act gat tct tcg ttt gac ggt ttt gat ctc Pro His Tyr Asp Met Ser Thr Asp Ser Ser Phe Asp Gly Phe Asp Leu 100 105 110	434										
tgg gg Trp	439										
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Met Met Leu Leu Ala Leu Phe Phe Leu Leu -15 -10											
agg att gct ttg gct agt caa ggt ctt ttg tgg ttc cat acc sat ttt Arg Ile Ala Leu Ala Ser Gln Gly Leu Leu Trp Phe His Thr Asn Phe -5 1 5 10	159										
aag gtt ttt gtt gtt tcy att tgt gtg aag act atc att ggg att tcg Lys Val Phe Val Val Ser Ile Cys Val Lys Thr Ile Ile Gly Ile Ser 15 20 25	207										
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											gtg Val	acc				103
ctg					tcg					cca	gga Gly			gcc		151
											gtt Val					199
											atc Ile 50				acc Thr	247
_							_	_			ggc Gly					295
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											ctt Leu 130					487
_	aca Thr															493
<212 <212	0 > 6: 1 > 1: 2 > Di 3 > Ho	0 8 N.A.	sapi	ens												
	1 > C	os 61	79													
<222	2 > 3 ( 3 > V( s)	on He	eptio 0 eijno 10.0 LFLL	e mai	0038		7									
	0> 6 cagt		tcca	gctc	cc a	aata	tagai	t ati	1		agg ( Arg )			Phe 1		53
									ggt	ctc	gct Ala			ctc	agg	101
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		agc					tca	gcc Ala		a		_			,	180

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                                                                       55
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                                                          -15
tte etc etc etg gtg tea get ecc aga tgg gte etg tet eag gtg eag
                                                                      103
Phe Leu Leu Val Ser Ala Pro Arg Trp Val Leu Ser Gln Val Gln
            -10
                                -5
cta cag gag tcg ggc cca gga ctg gtg aag cct tcg ggg agg ctg tcc
                                                                      151
Leu Gln Glu Ser Gly Pro Gly Leu Val Lys Pro Ser Gly Arg Leu Ser
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ctc gcc tgc gat gtg gtg gaa ttg agt ccg ccg gcc ccc agg ggc ggg
                                                                      199
Leu Ala Cys Asp Val Val Glu Leu Ser Pro Pro Ala Pro Arg Gly Gly
                    25
                                        30
tot goa gtg cat ctc aga aat ctt toa toa tgg gag ccc cac cta caa
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Ser Ala Val His Leu Arg Asn Leu Ser Ser Trp Glu Pro His Leu Gln
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ccc gtc tcg ggg
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Pro Val Ser Gly
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       Met Thr Tyr Phe Pro Leu Gly Arg Tyr Pro Val Met Gly Leu
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                                   -25
chg gat caa atg gta gtt gtg ttt tta ctt ct/ tta gtd tcc.aca ctt
                                                                       96
Leu Asp Gln Met Val Val Val Phe Leu Leu Leu Val Ser Thr Leu
                                -10
tot too gta gtg gtt tta ota gtt tgc att occ acc agc agt gta aaa
Ser Ser Val Val Leu Leu Val Cys Ile Pro Thr Ser Ser Val Lys
ttg ttc cct ttt cac cat atc cac acc aac tgg g
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Leu Phe Pro Phe His His Ile His Thr Asn Trp

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35

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tat t	ta :	act	atc	tta	ttc	tac	ctc	tct	ctc	tcc	tta	taa			tat	162
Tyr I																
tta c	ett	ttt	ttg	ctt	ttt	gct	tgg	cct	ggg							192
Leu I		Phe -5	Leu	Leu	Phe	Ala	Trp 1	Pro	Gly							
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agago	ctct	<b>9</b> 9 9	gagt	ctg	a co											52
						-20		a Trj	Thi	Pro	. Lei 15		ı Phe	e Let	ı Thr	
ctc c	ctc	ctc	cac	tgc	aca	999	tct	ctc	gcc	cag	ctt	gtg	ctg	act	caa	100
Leu I	Leu	Leu	His	Cys		Gly	Ser	Leu	Ala	_	Leu	Val	Leu	Thr 5	Gln	
-10 tcg (	~~~	tct	acc	tct	-5 acc	tcc	cta	gga	acc	l tra	atc	аад	ctc	-	tac	148
Ser I																
act o	cta	agc		aga	cac	aqc	aac		qqc	atc	qct	tgg	_	caq	caq	196
Thr I																
		25					30					35				
cag d																244
Gln 1	40	GIU	пåв	GIA	FLU	45	FIIG	neu	PIC!	шус	50	asii.	SET	voh	J.y	
agc :	cac														agč -	2.93
Ser 1	His	Met	Lys	Ala	_	Gly	Ile	Pro	Asp	_	Phe	Ser	Gly	Ser		
55 tct 9	~~~	<b>ac</b> +	~~~	~~~	60	ctc	tcc	atc	tcc	65	ctc	2			70 .	329
Ser (												<b>~</b>				363

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WO 99/53051 51 <211> 314 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 259..312 <221> sig_peptide <222> 259..300 <223> Von Heijne matrix score 10.3999996185303 seq PLALFFLLSVALA/IQ <400> 75 tagggtgaga gatggggatc tagttttatt cttctgcata tggatatcca gttttcccag taacatttat tqaaqaqact qqcctttccc caatqagtqt tcttggcacc tttqtcaaaa 120 qtcaqttqqc cgtagatatg tggattaatt tctgtgttcc ctgttttgtt ccattggcct 180 atqtqtctqt ttttatgaca gtaccaggtt gttttggtta ctacagcttt gtagtttact 240 ttqaqqtctq ttaqtqtg atg cct cta gct ttg ttc ttt ttg ctc agt gtt 291 Met Pro Leu Ala Leu Phe Phe Leu Leu Ser Val 314 gct ttg gct att cag ggt cag gg Ala Leu Ala Ile Gln Gly Gln 1 <210> 76

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cct qqq qcc tca gtg aag gtt tcc tgc aag gca tct gga tac acc ttc 202 Pro Gly Ala Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe 250 ace age cae tat atg cae tgg gtg ega cag gee eet gga caa ggg ett Thr Ser His Tyr Met His Trp Val Arg Gln Ala Pro Gly Gln Gly Leu 35 gag tog and oga atalanth had not gat agti gat acc act sag had dha 295 Glu Trp Met Gly Ile Ile Tyr Pro Asp Ser Asp Thr Thr Lys Tyr Xaa 50 55 346 cag aac ttc cag ggc aga gtc acc atg act agg gac acg tcc acg agc Gln Asn Phe Gln Gly Arg Val Thr Met Thr Arg Asp Thr Ser Thr Ser

65 70 aca gtc tac atg gag ctg agc agc ctg aca tct gac gac acg gcc gtg 394 Thr Val Tyr Met Glu Leu Ser Ser Leu Thr Ser Asp Asp Thr Ala Val

score 10.1999998092651 seq XLXLSVLLGXXXX/KX

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                            85
                                                 90
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Trp
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                                          -20
tte etc etc etg etg geg get ecc aga tgg gte etg tec eag etg
                                                                        99
Phe Leu Leu Leu Val Ala Ala Pro Arg Trp Val Leu Ser Gln Leu
                -10
                                     -5
cag ctt cag gag tcg ggc cca gga ctg gtg aag gct tcg gag acc ctg
                                                                       147
Gln Leu Gln Glu Ser Gly Pro Gly Leu Val Lys Ala Ser Glu Thr Leu
                             10
tcc ctc gcc tgc agt gtc tct ggt gac tcc atc agc agt ggt aat tat
                                                                       195
Ser Leu Ala Cys Ser Val Ser Gly Asp Ser Ile Ser Ser Gly Asn Tyr
                         25
                                             30
tac tgg ggc tgg atc cgg cag ccc cca ggg aag gga ctg cag tgg ctt
                                                                       243
Tyr Trp Gly Trp Ile Arg Gln Pro Pro Gly Lys Gly Leu Gln Trp Leu
                                         45
ggg agt ctt tgg aat cgt ggc ggt ccg caa tac aay hcc tcc ctc aag
                                                                       291
Gly Ser Leu Trp Asn Arg Gly Gly Pro Gln Tyr Asn Xaa Ser Leu Lys
                                     60
aat cga gtc acc gtg tcc gta gac acg tcc acg aat cat ttc ttt ctg
                                                                      339
Asn Arg Val Thr Val Ser Val Asp Thr Ser Thr Asn His Phe Phe Leu
aga ctg aat tcc gtg aay vgh gga cac ggc aat tta tta ctg tgc gcg a
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-5 1 5 ctc cga atc tct gta gcc gat g Leu Arg Ile Ser Val Ala Asp 10 15	121
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Phe Phe Leu Leu Phe Phe Cys Phe Val Phe Cys Leu Arg Gly Gln Gly -10 -5 1	149
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gct ttt tta aaa ggt gtc cag tgt gag gtg cag ttg ttg gag tct ggg Ala Phe Leu Lys Gly Val Gln Cys Glu Val Gln Leu Leu Glu Ser Gly	160
gga ggc ttg gtc cag cct ggg ggg tcc ctg aga ctc tca tgt gca gcc Gly Gly Leu Val Gln Pro Gly Gly Ser Leu Arg Leu Ser Cys Ala Ala	208

10 15 20	
tcc gga ttc acc ttt agc tcc tat gcc atg ctc tgg gtc cgc cag gct Ser Gly Phe Thr Phe Ser Ser Tyr Ala Met Leu Trp Val Arg Gln Ala 25 30 35 40	256
cca ggt aag ggg ctg gag tgg gtc tca ggt att agt gct ggt gct gat Pro Gly Lys Gly Leu Glu Trp Val Ser Gly Ile Ser Ala Gly Ala Asp 45 50 55	304
gat aca tat gat gca gac tcc gtg aag ggc cgg ttc acc att tcc aga Asp Thr Tyr Asp Ala Asp Ser Val Lys Gly Arg Phe Thr Ile Ser Arg 60 65 70	352
gac gat tcc aag aaa atc cta tat cta caa atg aac agc ctg aga gcc Asp Asp Ser Lys Lys Ile Leu Tyr Leu Gln Met Asn Ser Leu Arg Ala 75 80 85	400
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Arg Leu Thr Val Val Leu Gly Leu Leu Val Leu Phe Leu Thr Cys Tyr -15 -10 -5	
· ·	151
-15 -10 -5 gca gac gac aaa cca gac aag cca gac gac	151
gca gac gac aaa cca gac aag cca gac gac	
gca gac gac aaa cca gac aag cca gac gac	199
-15	199

score 10

seq LLLLQALPSPLSA/RA

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ctc ctg ctc ctc cag gcg ctg ccc agc ccc ttg tca gcc agg gct gaa 161 Leu Leu Leu Gln Ala Leu Pro Ser Pro Leu Ser Ala Arg Ala Glu -10 -5 1
ccc ccg cag gat aag gaa gcc tgt gtg ggt acc aac aat caa agc tac 209 Pro Pro Gln Asp Lys Glu Ala Cys Val Gly Thr Asn Asn Gln Ser Tyr 5 10 15
atc tgt gac aca gga cac tgc tgt gga cag tct cag tgc tgy aac tac 257  Ile Cys Asp Thr Gly His Cys Cys Gly Gln Ser Gln Cys Cys Asn Tyr  20 25 30 35
tac tat gaa ctc tgg tgg ttc tgg ctg gtg tgg acc atc atc atc atc  Tyr Tyr Glu Leu Trp Trp Phe Trp Leu Val Trp Thr Ile Ile Ile  40  45  50
ctg agc tgc tgc tgt gtt tgc cac cac cgc cga gcc aag cac cgc ctt  Leu Ser Cys Cys Cys Val Cys His His Arg Arg Ala Lys His Arg Leu  55 60 65
cag gcc cag cag cgg caa cat gaa atc aac ctg atc gct tac cga g 399 Gln Ala Gln Gln Arg Gln His Glu Ile Asn Leu Ile Ala Tyr Arg 70 75 80
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taggtataac acattaagaa aaagttatct tcattggata gaattga atg gtg gtc 296  Met Val Val  -25
gct gat agg aat agg gcg tcc tct agc tct tat ctc tgt ctc tta ctc  Ala Asp Arg Asn Arg Ala Ser Ser Ser Ser Tyr Leu Cys Leu Leu  -20  -15  -10
ttt tct ctt tct ctt ttt ctc tgt cat gag act gtg tgt gac agg gcc  Phe Ser Leu Ser Leu Phe Leu Cys His Glu Thr Val Cys Asp Arg Ala  -5  1  5
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-35 -30 -25	
aac agg aga acc aag acc aaa act tta ttg tct ctg ctt tca ttt ctt 14	9
Asn Arg Arg Thr Lys Thr Leu Leu Ser Leu Leu Ser Phe Leu -20 -15 -10	
gat gaa acc tot gga cta agc aca cat ott cot tgt tta tot otc tca 19	7
Asp Glu Thr Ser Gly Leu Ser Thr His Leu Pro Cys Leu Ser Leu Ser -5 1 5 10	
aag gag tgt gga gtg ctt cat ctg gac atc cac ggg aag aag gaa gac 24.	5
Lys Glu Cys Gly Val Leu His Leu Asp Ile His Gly Lys Lys Glu Asp 15 20 25	
atg aga gat gag gtc ttg ctg gcc ttg aac tyc tgc acc cac agg 29	O
Met Arg Asp Glu Val Leu Leu Ala Leu Asn Xaa Cys Thr His Arg	_
30 35 40	
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Met Lys Ser Phe Ser	•
-15	
egg atc etc tte etc gte tte etc etc gee gge etg agg tec aag gee 16	2
Arg Ile Leu Phe Leu Val Phe Leu Leu Ala Gly Leu Arg Ser Lys Ala	
-10 -5 1	
get eec tea gee eet etg eet ttg gge tgt gge ttt eeg gae atg gee 21	0
Ala Pro Ser Ala Pro Leu Pro Leu Gly Cys Gly Phe Pro Asp Met Ala	
5 10 15	
cac ccc tct gag act tcc cct ctg aag ggt gct tct gaa aat tcc aaa 25	8
His Pro Ser Glu Thr Ser Pro Leu Lys Gly Ala Ser Glu Asn Ser Lys	
20 25 30	_
cga gat cgc ctt aac cca gaa ttt cct ggg act cct tac cct gag cct	6
Arg Asp Arg Leu Asn Pro Glu Phe Pro Gly Thr Pro Tyr Pro Glu Pro 35 40 45 50	
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For manners of the second second

58

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<213> Homo sapiens

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aaa ccc ttc ttt Lys Pro Phe Phe	agc cct tca	cac atc gca	ctg aag aat at	
1	5	10		15
aag gat atg gaa Lys Asp Met Glu 20				<del>-</del> -
gat gat gat gag Asp Asp Asp Glu 35				
cat ttt ttt cca His Phe Phe Pro	Phe Asp Leu	ttt cca atg	tgt cca ttt gg Cys Pro Phe Gl	
50 tgc tat tca cga Cys Tyr Ser Arg	Val Val His	_	60	508
65	70			: ù
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Phe Leu Leu -10	Val Ala Ala	Pro Arg Trp -5	Ala Met Ser Gl	n Val Gln
cts cag gaa teg Leu Gln Glu Ser 5				
ctc acc tgc agt Leu Thr Cys Ser	gtc tct ggt	ggc tcc atg	gcc act agt ga	c tgg tgg 197
-cu in cys ser	var ser Giy	GIA SET MEC	via int ser We	- 115 115

30

245

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20

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<222> 15..92

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score 9.5

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<212> DNA

<213> Homo sapiens

Gly Leu Arg Val Ser

<220>

<221> CDS

<222> 118..315

<221> sig_peptide

<222> 118..306

63

<223> Von Heijne matrix

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Met Ala Asn Gly Thr Asn Ala Ser Ala Pro Tyr Tyr Ser Tyr Glu Tyr	165
-60 -55 -50	
tac ctg gac tat ctg gac ctc att ccc gtg gac gag aag aag ctg aaa	213
Tyr Leu Asp Tyr Leu Asp Leu Ile Pro Val Asp Glu Lys Lys Leu Lys	
-45 -40 -35	
gcc cac aaa cat tcc atc gtg atc gca ttc tgg gtg agc ctg gct gcc	261
Ala His Lys His Ser Ile Val Ile Ala Phe Trp Val Ser Leu Ala Ala	
-30 -25 -20	
ttc gtg gtg ctg ctc ttc ctc atc ttg ctc tac atg tcc tgg tcc gcs	309
Phe Val Val Leu Leu Phe Leu Ile Leu Leu Tyr Met Ser Trp Ser Ala	
-15 -10 , -5 1	215
tcc ccg Ser Pro	315
SEL PIO	
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c atg gac tgg acc tgg agg ttc ctc ttt gtg gtg gca gca gct aca ggt	60
Met Asp Trp Thr Trp Arg Phe Leu Phe Val Val Ala Ala Ala Thr Gly	109
-15 -10 -5	109
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gtc cag tcm cag gks cas ctg gwg cag tct ggg gct gag gtg aag aag Val Gln Ser Gln Xaa Xaa Leu Xaa Gln Ser Gly Ala Glu Val Lys Lys	109
gtc cag tcm cag gks cas ctg gwg cag tct ggg gct gag gtg aag aag Val Gln Ser Gln Xaa Xaa Leu Xaa Gln Ser Gly Ala Glu Val Lys Lys 1 5 10	109 157
gtc cag tcm cag gks cas ctg gwg cag tct ggg gct gag gtg aag aag Val Gln Ser Gln Xaa Xaa Leu Xaa Gln Ser Gly Ala Glu Val Lys Lys  1 5 10 cct ggg tcc tcg gtg aaa gtc tcc tgc arg gcy tct gga ggc atc ytc	109
gtc cag tcm cag gks cas ctg gwg cag tct ggg gct gag gtg aag aag Val Gln Ser Gln Xaa Xaa Leu Xaa Gln Ser Gly Ala Glu Val Lys Lys  1 5 10  cct ggg tcc tcg gtg aaa gtc tcc tgc arg gcy tct gga ggc atc ytc Pro Gly Ser Ser Val Lys Val Ser Cys Xaa Ala Ser Gly Gly Ile Xaa	109 157
gtc cag tcm cag gks cas ctg gwg cag tct ggg gct gag gtg aag aag Val Gln Ser Gln Xaa Xaa Leu Xaa Gln Ser Gly Ala Glu Val Lys Lys  1 5 10  cct ggg tcc tcg gtg aaa gtc tcc tgc arg gcy tct gga ggc atc ytc Pro Gly Ser Ser Val Lys Val Ser Cys Xaa Ala Ser Gly Gly Ile Xaa 15 20 25	109 157 205
gtc cag tcm cag gks cas ctg gwg cag tct ggg gct gag gtg aag aag Val Gln Ser Gln Xaa Xaa Leu Xaa Gln Ser Gly Ala Glu Val Lys Lys  1 5 10  cct ggg tcc tcg gtg aaa gtc tcc tgc arg gcy tct gga ggc atc ytc Pro Gly Ser Ser Val Lys Val Ser Cys Xaa Ala Ser Gly Gly Ile Xaa  15 20 25  agc asc tat agc ttc aac tgg gtg cgm cag gcc cct gga cag ggg ttt	109 157
gtc cag tcm cag gks cas ctg gwg cag tct ggg gct gag gtg aag aag Val Gln Ser Gln Xaa Xaa Leu Xaa Gln Ser Gly Ala Glu Val Lys Lys  1 5 10  cct ggg tcc tcg gtg aaa gtc tcc tgc arg gcy tct gga ggc atc ytc Pro Gly Ser Ser Val Lys Val Ser Cys Xaa Ala Ser Gly Gly Ile Xaa  15 20 25  agc asc tat agc ttc aac tgg gtg cgm cag gcc cct gga cag ggg ttt Ser Xaa Tyr Ser Phe Asn Trp Val Arg Gln Ala Pro Gly Gln Gly Phe	109 157 205
gtc cag tcm cag gks cas ctg gwg cag tct ggg gct gag gtg aag aag Val Gln Ser Gln Xaa Xaa Leu Xaa Gln Ser Gly Ala Glu Val Lys Lys  1 5 10  cct ggg tcc tcg gtg aaa gtc tcc tgc arg gcy tct gga ggc atc ytc Pro Gly Ser Ser Val Lys Val Ser Cys Xaa Ala Ser Gly Gly Ile Xaa 15 20 25  agc asc tat agc ttc aac tgg gtg cgm cag gcc cct gga cag ggg ttt Ser Xaa Tyr Ser Phe Asn Trp Val Arg Gln Ala Pro Gly Gln Gly Phe 30 35 40 45	109 157 205 253
gtc cag tcm cag gks cas ctg gwg cag tct ggg gct gag gtg aag aag Val Gln Ser Gln Xaa Xaa Leu Xaa Gln Ser Gly Ala Glu Val Lys Lys  1 5 10  cct ggg tcc tcg gtg aaa gtc tcc tgc arg gcy tct gga ggc atc ytc Pro Gly Ser Ser Val Lys Val Ser Cys Xaa Ala Ser Gly Gly Ile Xaa  15 20 25  agc asc tat agc ttc aac tgg gtg cgm cag gcc cct gga cag ggg ttt Ser Xaa Tyr Ser Phe Asn Trp Val Arg Gln Ala Pro Gly Gln Gly Phe 30 35 40 45  gag tgg ttg gga agg atc atc ccc atc ctc ggt ata aca aac tac gca	109 157 205
gtc cag tcm cag gks cas ctg gwg cag tct ggg gct gag gtg aag aag Val Gln Ser Gln Xaa Xaa Leu Xaa Gln Ser Gly Ala Glu Val Lys Lys  1 5 10  cct ggg tcc tcg gtg aaa gtc tcc tgc arg gcy tct gga ggc atc ytc Pro Gly Ser Ser Val Lys Val Ser Cys Xaa Ala Ser Gly Gly Ile Xaa 15 20 25  agc asc tat agc ttc aac tgg gtg cgm cag gcc cct gga cag ggg ttt Ser Xaa Tyr Ser Phe Asn Trp Val Arg Gln Ala Pro Gly Gln Gly Phe 30 35 40 45	109 157 205 253
gtc cag tcm cag gks cas ctg gwg cag tct ggg gct gag gtg aag aag Val Gln Ser Gln Xaa Xaa Leu Xaa Gln Ser Gly Ala Glu Val Lys Lys  1 5 10  cct ggg tcc tcg gtg aaa gtc tcc tgc arg gcy tct gga ggc atc ytc Pro Gly Ser Ser Val Lys Val Ser Cys Xaa Ala Ser Gly Gly Ile Xaa 15 20 25  agc asc tat agc ttc aac tgg gtg cgm cag gcc cct gga cag ggg ttt Ser Xaa Tyr Ser Phe Asn Trp Val Arg Gln Ala Pro Gly Gln Gly Phe 30 35 40 45  gag tgg ttg gga agg atc atc ccc atc ctc ggt ata aca aac tac gca Glu Trp Leu Gly Arg Ile Ile Pro Ile Leu Gly Ile Thr Asn Tyr Ala	109 157 205 253
gtc cag tcm cag gks cas ctg gwg cag tct ggg gct gag gtg aag aag Val Gln Ser Gln Xaa Xaa Leu Xaa Gln Ser Gly Ala Glu Val Lys Lys  1	109 157 205 253 301
gtc cag tcm cag gks cas ctg gwg cag tct ggg gct gag gtg aag aag Val Gln Ser Gln Xaa Xaa Leu Xaa Gln Ser Gly Ala Glu Val Lys Lys  1	109 157 205 253 301
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gtc cag tcm cag gks cas         ctg gwg cag tct ggg gct gag gtg aag aag           Val Gln Ser Gln Xaa Xaa Leu Xaa Gln Ser Gly Ala Glu Val Lys Lys           1         5           10           Cct ggg tcc tcg gtg aaa gtc tcc tgc arg gcy tct gga ggc atc ytc           Pro Gly Ser Ser Val Lys Val Ser Cys Xaa Ala Ser Gly Gly Ile Xaa           15         20           25           agc asc tat agc ttc aac tgg gtg cgm cag gcc cct gga cag ggg ttt           Ser Xaa Tyr Ser Phe Asn Trp Val Arg Gln Ala Pro Gly Gln Gly Phe           30         35           40         45           gag tgg ttg gga agg atc atc ccc atc ctc ggt ata aca aac tac gca           Glu Trp Leu Gly Arg Ile Ile Pro Ile Leu Gly Ile Thr Asn Tyr Ala           50         55           50           gag aag ttt cgg ggc aga ctc acg atc acc gtg gac aaa tcc acg cgt           Glu Lys Phe Arg Gly Arg Leu Thr Ile Thr Val Asp Lys Ser Thr Arg           65         70           75           gtt gtt tac atg gag cag agc agc agc agc ctg aca tct gcg gac acg gcc gta           Val Val Tyr Met Glu Gln Ser Ser Leu Thr Ser Ala Asp Thr Ala Val           80         85           90           tat tat tgt gcg aaa ccg acc act atg act tcg gag cta act tat	109 157 205 253 301 349
gtc cag tcm cag gks cas ctg gwg cag tct ggg gct gag gtg aag aag           Val Gln Ser Gln Xaa Xaa Leu Xaa Gln Ser Gly Ala Glu Val Lys Lys           1         5         10           cct ggg tcc tcg gtg aaa gtc tcc tgc arg gcy tct gga ggc atc ytc           Pro Gly Ser Ser Val Lys Val Ser Cys Xaa Ala Ser Gly Gly Ile Xaa         25           agc asc tat agc ttc aac tgg gtg cgm cag gcc cct gga cag ggg ttt         25           agc asc tat agc ttc aac tgg gtg cgm cag gcc cct gga cag ggg ttt         35         40         45           gag tgg ttg gga agg atc atc ccc atc ctc ggt ata aca aac tac gca         35         40         45           gag tgg ttg gga agg atc atc acc ccc atc ctc ggt ata aca aac tac gca         35         50         50           gag aag ttt cgg ggc aga ctc acg atc acc gtg gac aaa tcc acg cgt         36         36         36         36         36         36         36         36         36         36         36         36         36         36         36         36         36         36         36         36         36         36         36         36         36         36         36         36         36         36         36         36         36         36         36         36         36         36         36         36         36         36         3	109 157 205 253 301 349
gtc cag tcm cag gks cas         ctg gwg cag tct ggg gct gag gtg aag aag           Val Gln Ser Gln Xaa Xaa Leu Xaa Gln Ser Gly Ala Glu Val Lys Lys           1         5           10           Cct ggg tcc tcg gtg aaa gtc tcc tgc arg gcy tct gga ggc atc ytc           Pro Gly Ser Ser Val Lys Val Ser Cys Xaa Ala Ser Gly Gly Ile Xaa           15         20           25           agc asc tat agc ttc aac tgg gtg cgm cag gcc cct gga cag ggg ttt           Ser Xaa Tyr Ser Phe Asn Trp Val Arg Gln Ala Pro Gly Gln Gly Phe           30         35           40         45           gag tgg ttg gga agg atc atc ccc atc ctc ggt ata aca aac tac gca           Glu Trp Leu Gly Arg Ile Ile Pro Ile Leu Gly Ile Thr Asn Tyr Ala           50         55           50           gag aag ttt cgg ggc aga ctc acg atc acc gtg gac aaa tcc acg cgt           Glu Lys Phe Arg Gly Arg Leu Thr Ile Thr Val Asp Lys Ser Thr Arg           65         70           75           gtt gtt tac atg gag cag agc agc agc agc ctg aca tct gcg gac acg gcc gta           Val Val Tyr Met Glu Gln Ser Ser Leu Thr Ser Ala Asp Thr Ala Val           80         85           90           tat tat tgt gcg aaa ccg acc act atg act tcg gag cta act tat	109 157 205 253 301 349

64 cag wct aca cta tgg 460 Gln Xaa Thr Leu Trp <210> 98 <211> 230 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 140..229 <221> sig_peptide <222> 140..205 <223> Von Heijne matrix score 9.39999961853027 seq LLLLSAFTSQTVS/GQ <400> 98 aacagaacaa tatcaaatag ctaacttcac ccccaaccac agtccttgct gttggcattt 60 actcaactag tctttaattc ctgttttgac aaactttata aggtgctaca agacagatga 120 tttttcacca tctaccata atg tgg aac aga tat ttt gtc ttc tat ctc ctg 172 Met Trp Asn Arg Tyr Phe Val Phe Tyr Leu Leu -20 ctt ttg tca gcg ttt acg agt caa aca gta tcc gga caa aga aag aaa 220 Leu Leu Ser Ala Phe Thr Ser Gln Thr Val Ser Gly Gln Arg Lys Lys -10 gga ccc cgg g 230 Gly Pro Arg <210> 99 <211> 467 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 40..465 <221> sig_peptide <222> 40..96 <223> Von Heijne matrix score 9.39999961853027 seq FLLLVAAPRWVLS/QL aaatactttc tgagagccct ggacctcctg tgcaagaac atg aaa cac ctg ggg 54 Met Lys His Leu Gly -15 ttc ttc ctc ctg ctg gtg gca gct ccc aga tgg gtc ctg tcc cag ctg 102 Phe Phe Leu Leu Val Ala Ala Pro Arg Trp Val Leu Ser Gln Leu -10 cag etc cag gag tec gge tea gga etg gag aag eet tee cag acc etg 150 Gln Leu Gln Glu Ser Gly Ser Gly Leu Glu Lys Pro Ser Gln Thr Leu 20 tee etc ace tge tet gte tet ggt gge tee ate agt agt gat gat ttg 198 Ser Leu Thr Cys Ser Val Ser Gly Gly Ser Ile Ser Ser Asp Asp Leu 25 teg tgg age tgg ate ega eag eeg eea ggg aag gge etg gag tgg att 246 Ser Trp Ser Trp Ile Arg Gln Pro Pro Gly Lys Gly Leu Glu Trp Ile 40 45

ggc tac att tat caa aat gag agg acc ctc tac aac ccg tcc ctc aag

Gly Tyr Ile Tyr Gln Asn Glu Arg Thr Leu Tyr Asn Pro Ser Leu Lys 60 agt ega gee gee att tea gtg gae agg tee aag aac eag tte tee etg 342 Ser Arg Ala Ala Ile Ser Val Asp Arg Ser Lys Asn Gln Phe Ser Leu aaa ctg acc tct gtg acc gcc gcg gac atg gcc gta tat tac tgt gcc 390 Lvs Leu Thr Ser Val Thr Ala Ala Asp Met Ala Val Tyr Tyr Cys Ala acc agt gtc atg awt tcc ttt ggg ggc gtt ctc gtc cct aat ctg ttt 438 Thr Ser Val Met Xaa Ser Phe Gly Gly Val Leu Val Pro Asn Leu Phe 110 105 ttg act act ggg gcc agg gaa tct cgt ca 467 Leu Thr Thr Gly Ala Arg Glu Ser Arg 115 <210> 100 <211> 504 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 39..503 <221> sig peptide <222> 39..95 <223> Von Heijne matrix score 9.30000019073486 seq FLLLVAGPRWVLS/QV <400> 100 56 aatactttct gagagtcctg gacctcctgt gcaagaac atg aaa cac ctg tgg ttc Met Lys His Leu Trp Phe tte etc etg etg gtg gea ggt eec aga tgg gte etg tee eag gtg eag 104 Phe Leu Leu Val Ala Gly Pro Arg Trp Val Leu Ser Gln Val Gln -5 ctq sdk qaq tcq ggc cca aga ctg gtg aag cct tca cag acc ctg tcc 152 Leu Xaa Glu Ser Gly Pro Arg Leu Val Lys Pro Ser Gln Thr Leu Ser ctc acc tgc act gta tct ggg gcc tcc gtc agc agt cgt ggg tac tat 200 Leu Thr Cys Thr Val Ser Gly Ala Ser Val Ser Ser Arg Gly Tyr Tyr tgg acc tgg atc cgc cag ctc cca ggg aag ggc ctg gag tgg att ggc 248 Trp Thr Trp Ile Arg Gln Leu Pro Gly Lys Gly Leu Glu Trp Ile Gly 40 tac atc tgk tac act ggg agc acc ttc tac aac ccg tcc ctc aag agt 296 Tyr Ile Xaa Tyr Thr Gly Ser Thr Phe Tyr Asn Pro Ser Leu Lys Ser 60 cga tta acc ata tca ata gac acg tct aag aat cag ttc tcc ctg aac 344 Arg Leu Thr Ile Ser Ile Asp Thr Ser Lys Asn Gln Phe Ser Leu Asn 70 75 ctg agg tct gtg act acc gcg gac acg gcc gtc tat tac tgt gcg aga 392 Leu Arg Ser Val Thr Thr Ala Asp Thr Ala Val Tyr Tyr Cys Ala Arg 90 95 gad cat ttp gat ctt cta ttp gac ccc tgg ggc dag gga acc ckg gto-440

Asp His Phe Asp Leu Leu Phe Asp Pro Trp Gly Gin Gly Thr Leu Val

ace gto tee tet gem tee ace aag gge eea teg gto tto eec etg gea

Thr Val Ser Ser Ala Ser Thr Lys Gly Pro Ser Val Phe Pro Leu Ala

110

125

488

504

130

105

120

scc tcc tcc aag agc a Xaa Ser Ser Lys Ser

135

His Ala

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                                                                       60
tattetacct etceettget etttecagae caggettggg acataacact aacaccettt
                                                                      120
tcctttcatt tcatctcttg tccttcagtc attcctaaac attgacaybc attgacttcc
                                                                      180
ttggctctgg ccatagtcct ttctcccttt cccctctggg gcatcaaata gtgattacag
                                                                      240
tatccacagg g atg gca tat gcc att tca cca ttt cac agt tcc tgg aat
                                                                      290
             Met Ala Tyr Ala Ile Ser Pro Phe His Ser Ser Trp Asn
                    -40
                                         -35
cca ctt ttc act tct cat aaa gct tca gca agc cat tct cat ctt ggg
                                                                      338
Pro Leu Phe Thr Ser His Lys Ala Ser Ala Ser His Ser His Leu Gly
                -25
                                    -20
ttg ctt gtt tgc cta ttt gct gtt aca tcc att ctc tgc tcc tca
                                                                      383
Leu Leu Val Cys Leu Phe Ala Val Thr Ser Ile Leu Cys Ser Ser
            -10
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                                                                       53
                                Met Ser Pro Val Leu Leu Ala
                                -15
                                                     -10
cto cto ago the atc etc cca etg cca ggm agt gca rud get gas tek-
                                                                      101
Leu Leu Gly Phe Ile Leu Pro Leu Pro Gly Ser Ala Xaa Ala Xaa Ser
        -5
                            1
gcc agt ttg gga cag ttc agc atg tgt gga agg tgt ccg acm tgc ccc
                                                                      149
Ala Ser Leu Gly Gln Phe Ser Met Cys Gly Arg Cys Pro Thr Cys Pro
                    15
                                                             25
                                        20
ggc aat gga ccc cta aga aca cca gct gcg aca sgg vtt rgg gtg cca
                                                                      197
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Gly Asn Gly Pro Leu Arg Thr Pro Ala Ala Thr Xaa Xaa Xaa Val Pro

68

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35
                                                         40
gga cac gtt gat gc
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Gly His Val Asp
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                                                                        52
                                Met Ala Thr Ala Met Asp Trp Leu
                                            -20
ccg tgg tct tta ctg ctt ttc tcc ctg atg tgt gaa aca agc gcc ttc
                                                                       100
Pro Trp Ser Leu Leu Leu Phe Ser Leu Met Cys Glu Thr Ser Ala Phe
                    -10
                                         -5
tat gtg cct ggg gtc gcg cct atc aac ttc cac cag aac gat ccc gta
                                                                       148
Tyr Val Pro Gly Val Ala Pro Ile Asn Phe His Gln Asn Asp Pro Val
                                 10
gaa atc aag gct gtg aag ctc acc agc tct cga acc cag cta cct tat
                                                                       196
Glu Ile Lys Ala Val Lys Leu Thr Ser Ser Arg Thr Gln Leu Pro Tyr
                             25
gaa tac tat tca ctg ccc ttc tgc cag ccc agc aag ata acc tac aag
                                                                       244
Glu Tyr Tyr Ser Leu Pro Phe Cys Gln Pro Ser Lys Ile Thr Tyr Lys
                         40
                                             45
gca gag aat ctg gga gag gtg ctg aga ggg gac cgg att gtc aac acc
                                                                       292
Ala Glu Asn Leu Gly Glu Val Leu Arg Gly Asp Arg Ile Val Asn Thr
                    55
                                         60
cct ttc cag gtt ctc atg aac agc gag aag aag tgt gaa gtt ctg tgc
                                                                       340
Pro Phe Gln Val Leu Met Asn Ser Glu Lys Lys Cys Glu Val Leu Cys
                                     75
age cag tee aac aag eea gtg ace etg aca gtg gag cag age ega etc
                                                                       388
Ser Gln Ser Asn Lys Pro Val Thr Leu Thr Val Glu Gln Ser Arg Leu
gtg gcc gag cgg atc aca gaa gac tac tac gtc cac ctc att gct gac
                                                                       436
Val Ala Glu Arg Ile Thr Glu Asp Tyr Tyr Val His Leu Ile Ala Asp
                             105
aac ctg cct gtg gcc acc ggc tgg agc tct act cca acc gag aca gcg
                                                                       484
Asn Leu Pro Val Ala Thr Gly Trp Ser Ser Thr Pro Thr Glu Thr Ala
    115
atg aca ag
                                                                       492
Met Thr
740
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seq WIFLLAILKGVQC/EV

WO 99/53051 PCT/IB99/00712

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71

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Ser Leu Ala Leu Leu Ala Ser Ser Pro Ile Ala Ala Xaa Pro

58

106

154

202

250

298

346

394

441

72 <210> 112 <211> 441 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 59..439 <221> sig_peptide <222> 59..115 <223> Von Heijne matrix score 8.89999961853027 seq ILLLVAAATDASS/QM <400> 112 atcacccatc aaccacatcc ctcctctaga gagtcccctg aaagcacagc tcctcacc atg gac tgg acc tgg aga atc ctc ctc ttg gtg gca gca gcc aca gat Met Asp Trp Trp Arg Ile Leu Leu Leu Val Ala Ala Ala Thr Asp -15 -10 gcc tcc tcc cag atg cag ctg ttg cag tct ggg cct gaa gtg aag aag Ala Ser Ser Gln Met Gln Leu Leu Gln Ser Gly Pro Glu Val Lys Lys act ggg tee tea gtg aaa ett tee tge aeg gee tee gge gae aee ete Thr Gly Ser Ser Val Lys Leu Ser Cys Thr Ala Ser Gly Asp Thr Leu 15 gcc tac cac tac ctg cac tgg gtg cga cag gcc ccc gga caa gcg ctt Ala Tyr His Tyr Leu His Trp Val Arg Gln Ala Pro Gly Gln Ala Leu 35 40 gag tgg atg gga tgg atc aca cct ttc agt gga gac acc aac ttc gca Glu Trp Met Gly Trp Ile Thr Pro Phe Ser Gly Asp Thr Asn Phe Ala 50 55 cag cga ttc cag gac aga ctc acc ttc acc agg gac agg tct atg agc Gln Arg Phe Gln Asp Arg Leu Thr Phe Thr Arg Asp Arg Ser Met Ser 70 aca gtc tac atg acc ctg acc agc ctg ata tct gaa gac aca gcc atg Thr Val Tyr Met Thr Leu Thr Ser Leu Ile Ser Glu Asp Thr Ala Met 80 85 90 tat tac tgt gcc act gat gga cgt cgc acc aac cgt ctt ttt gaa ca Tyr Tyr Cys Ala Thr Asp Gly Arg Arg Thr Asn Arg Leu Phe Glu 95 100 105 <210> 113 <211> 369 <212> DNA

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tca gtc agg gat gcc tac tgg aag act ggt agc tgc cca cct cca ttt Ser Val Arg Asp Ala Tyr Trp Lys Thr Gly Ser Cys Pro Pro Pro Phe 5 10 15	271
ctc cat gtg tct acc ttc nnn kkt aaa ctt acc ttc tcc act aag ggc Leu His Val Ser Thr Phe Xaa Xaa Lys Leu Thr Phe Ser Thr Lys Gly 20 25 30	319
aac ctt ctg cat tcc att cct ctc tct tcc ccc tta gcc tgt gtt ctt Asn Leu Leu His Ser Ile Pro Leu Ser Ser Pro Leu Ala Cys Val Leu 35 40 45 50	367
ag	369
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agt ccg agt cat ttc tca gaa ggt ttt gat agg tgg gcc tta gag gag Ser Pro Ser His Phe Ser Glu Gly Phe Asp Arg Trp Ala Leu Glu Glu -80 -75 -70 -65	100
	148
ctc gtg gtc agc att gca ctt aac ctc cag aag tac tgc cac atc cgc Leu Val Val Ser Ile Ala Leu Asn Leu Gln Lys Tyr Cys His Ile Arg -45 -40 -35	196
ctg gca ggc tcc aag gat ccc cgg gcc tat ttc aag acc aag aca tgg Leu Ala Gly Ser Lys Asp Pro Arg Ala Tyr Phe Lys Thr Lyc The Erp -30 -25 -20	244
· ·	292
ton tac gcc ttc gcg ccg ctg tca ctc atc gtg ccc ctc agc Ser Tyr Ala Phe Ala Pro Leu Ser Leu Ile Val Pro Leu Ser 1 5 10	334

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											aat Asn			_		103
	Phe				Pro					Tyr	aca Thr				Gly	151
cat	1	***			5					10					15	100
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agg gtt gaa tat tct cca gtg tct gtc tgc ttt tta tta ctt tcc gtt  Arg Val Glu Tyr Ser Pro Val Ser Val Cys Phe Leu Leu Ser Val  -20  -15  -25  -25  -25  -25  -25  -25  -25	221
gcc ttc aat cag ttg gtt ttt gct ttg tat cca ata caa gct acw btc Ala Phe Asn Gln Leu Val Phe Ala Leu Tyr Pro Ile Gln Ala Thr Xaa -5 1 5	269
tgt ttc tct dda gtt tct ctc cct ttc ccc gct ca Cys Phe Ser Xaa Val Ser Leu Pro Phe Pro Ala 10 15	304
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ctg tgg ccc ttt gcw ctt gct ctc tta aag acc c Leu Trp Pro Phe Ala Leu Ala Leu Leu Lys Thr 1 5	145
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205

386

att ggv ctg aag ttt tct ttt ttg ttg ttg cat ctc tgc cag gtt ttg 222 Ile Gly Leu Lys Phe Ser Phe Leu Leu His Leu Cys Gln Val Leu -10

cta tca aga cga gct ggt acc att cct act gaa aca att cca aaa aaa 270 Leu Ser Arg Arg Ala Gly Thr Ile Pro Thr Glu Thr Ile Pro Lys Lys

10 288 ttg agg agg aga gac ggg

Leu Arg Arg Arg Asp Gly 20

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<211> 386

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-15

-20 157 ctc ctg atg ggt ttc ctg atg gtc tgc ctg ggg gcc ttc ttc att tcc Leu Leu Met Gly Phe Leu Met Val Cys Leu Gly Ala Phe Phe Ile Ser

tgg ggc tcc ata ttc gac tgt cag ggg agc ctg att gcg gcc tat ttg Trp Gly Ser Ile Phe Asp Cys Gln Gly Ser Leu Ile Ala Ala Tyr Leu

10 15 20

ctt ctg cct ctg ggg ttt gtg atc ctt ctg agt gga att ttc tgg agc 253 Leu Leu Pro Leu Gly Phe Val Ile Leu Leu Ser Gly Ile Phe Trp Ser 30

aac tat cgc cag gtg act gaa agc aaa gga gtg ttg agg cac atg ctc 301 Asn Tyr Arg Gln Val Thr Glu Ser Lys Gly Val Leu Arg His Met Leu

45

cga caa cac ctt gct cat gyg gcc ctg ccc gtg gcc aca gta gac agt 349 Arg Gln His Leu Ala His Gly Ala Leu Pro Val Ala Thr Val Asp Ser

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Ala Ala Leu Leu Lys Ile Met Cys Lys Gln Leu Leu 75

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                                         Met Lys Val Glu Gly Glu Glu
   aag ctg tat cga ttg ttg aga tct ggc gac ttg ttt aaa ttt cat cag
                                                                         102
   Lys Leu Tyr Arg Leu Leu Arg Ser Gly Asp Leu Phe Lys Phe His Gln
                                -30
           -35
   cct cac ttc tat gaa ctc tca ggc ctc acg tgt acc agc tct ctg ctc
                                                                         150
   Pro His Phe Tyr Glu Leu Ser Gly Leu Thr Cys Thr Ser Ser Leu Leu
                            -15
                                                -10
                                                                         190
   tcc ttt gcc ttg gga cgt tcc atc cct gga agt ttc cca g
   Ser Phe Ala Leu Gly Arg Ser Ile Pro Gly Ser Phe Pro
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         seq LLLFSGAVALIQT/WA
   <400> 122
                                                                          52
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                                       Met Glu Ser Arg Thr Leu Leu
                                                       -15
   ctg ctg ttc tcg gga gcc gtg gcc ctg atc cag acc tgg gca ggt gag
                                                                         100
   Leu Leu Phe Ser Gly Ala Val Ala Leu Ile Gln Thr Trp Ala Gly Glu
            -10
                                -5
   tgc ggg gtc ggg agg gaa aag gcc tct gcg gga agg agc gag ggg ccc
                                                                         148
   Cys Gly Val Gly Arg Glu Lys Ala Ser Ala Gly Arg Ser Glu Gly Pro
                       10
                                            15
   gcc cgg agg agt aaa tct gca cat ata kbt aat tac aga tta caa tta
                                                                         196
   Ala Arg Arg Ser Lys Ser Ala His Ile Xaa Asn Tyr Arg Leu Gln Leu
                   25
saa toa agg cag ggg
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   Gln Ser Arg Gln Gly
               40
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<210> 123

<211> 353

<212> DNA

<213> Homo sapiens

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      seg SVPLLCFWSLCYC/FA
<221> misc feature
<222> 187
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                                                                       120
tetetgeete eagteeecea geecetggee gagagaaggg tettaeegge egggattget
                                                                       180
ggaaacncaa gaggtggttt ttgtttttta aaacttctgt ttcttgggag ggggtgtggc
                                                                       240
ggggcagg atg age aac tee gtt eet etg ete tgt tte tgg age ete tge
                                                                      290
         Met Ser Asn Ser Val Pro Leu Leu Cys Phe Trp Ser Leu Cys
             -15
                                 -10
tat tgc ttt gct gcg ggg agc ccc gta cct ttt ggt cca gag gga cgg
                                                                       338
Tyr Cys Phe Ala Ala Gly Ser Pro Val Pro Phe Gly Pro Glu Gly Arg
                                             10
                                                                       353
ctg gaa gat aag ctc
Leu Glu Asp Lys Leu
<210> 124
<211> 249
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      seg PWTILLFAAGSLA/IP
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                                                                       60
cagceggatt teccagecaa aegeagagag ag atg eee tgg ace ate ttg etc
                                                                       113
                                    Met Pro Trp Thr Ile Leu Leu
                                                     -10
ttt gca gct ggc tcc ttg gcg atc cca gca cca tcc atc cgg gtg gtg
                                                                       161
Phe Ala Ala Gly Ser Leu Ala Ile Pro Ala Pro Ser Ile Arg Val Val
                            1.
                                                                       209
ccc ccg tac cca agc agc caa gag gac ccc atc cac atc gca tgc atg
Pro Pro Tyr Pro Ser Ser Glu Glu Asp Pro I'le His I'le Ala Cys Met
                                                             25
gcc gct ggg aac ttc ccg ggg gcg aat ttc aca ctg tat c
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Ala Ala Gly Asn Phe Pro Gly Ala Asn Phe Thr Leu Tyr
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<210> 125 <211> 375

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-5

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ccctctgtcc ccgcggctgg gtctcgtctg ctccggttcc tgggctccta attcttggtc
                                                                      120
cagettette caggteagtg tgegggeett ceaegetgee ageggaacae tgga atg
                                                                      177
geg gaa ggg gaa egg gte tge geg tet gtk gtt eee age get etg ega
                                                                      225
Ala Glu Gly Glu Arg Val Cys Ala Ser Val Val Pro Ser Ala Leu Arg
                                -55
                                                                      273
acg ctg aaa agg agg agc aac ctg tcc aga atc ccc gca gga cag gaa
Thr Leu Lys Arg Arg Ser Asn Leu Ser Arg Ile Pro Ala Gly Gln Glu
                            -40
                                                 -35
aag gag ggg aaa tot oga oat gtt got ooc oot tit ogo tit tito oot
                                                                      321
Lys Glu Gly Lys Ser Arg His Val Ala Pro Pro Phe Arg Phe Pro
                        -25
                                             -20
ttt tcc ggt ttt ttg ttt ttt ggt ttt ctt ttt ccc gtc ttt tct ttc
                                                                      369
Phe Ser Gly Phe Leu Phe Gly Phe Leu Phe Pro Val Phe Ser Phe
                    -10
                                         -5
ccc tcc
                                                                      375
Pro Ser
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      seg MFCLAAILASASA/QR
<221> misc_feature
<222> 404
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                                                                      120
ggtataagta couttootto ttotytogtt maccaconco aggogagaaa actatgoooc
cgtgaaagtc cccactctgt ttcggttggg gaatactgga qcttaacctc ttggaggggg
                                                                      180
ttgttccata ccaagggtcc ttccgtaggt atttctaatg gg atg ttc tgc ctg
                                                                      234
                                                Met Phe Cys Leu
gca gca att tta gcc tca gca tct gcc caa cgg ttt cct tct gcc ttt
                                                                      282
Ala Ala Ile Leu Ala Ser Ala Ser Ala Gln Arg Phe Pro Ser Ala Phe
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tct cct tca cct tty yga t Ser Pro Ser Pro Phe Xaa T		
ttg ggt ttt trc act gtg t Leu Gly Phe Xaa Thr Val C	gy art aac tcc ata att	tcc ttg tgg tat 378
tta ayg ggr gtt ccc cca g Leu Xaa Gly Val Pro Pro G		
tgc agc atg gg Cys Ser Met		437
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	-15	-10
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aag tat ttc cac act tcc of Lys Tyr Phe His Thr Ser 10		
ttc atc tct gtg ggc tac g Phe Ile Ser Val Gly Tyr 1		
cgg gag aca cgg agc gcc a Arg Glu Thr Arg Ser Ala a 45		

ctg cgg acg ctg cgc ggc tac tac aat cag agc gag gcc ggg tct cam

Leu Arg Thr Leu Arg Gly Tyr Tyr Asn Gln Ser Glu Ala Gly Ser Xaa

70

65

293

304

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acc ctg cag tg Thr Leu Gln 75

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Ala P	tc gct he Ala 20														. 103
gag gg Glu G -5	gc gcc ly Ala	ctc Leu	ggc Gly	gag Glu 1	gag Glu	gct Ala	gca Ala	agt Ser 5	gcc Ala	gca Ala	gcc Ala	cag Gln	ggc Gly 10	cgc Arg	151
_	tg gct eu Ala					ca									174
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_	tt gcc le Ala	_													219
Phe L	tt tct eu Ser	Phe	Pro -5	Tyr	Thr	Leu	Cys	Ile 1	Leu	Tyr	Arg	Val 5	Lys	Ser	267
tat a Tyr T	ca ccc hr Pro	acg Thr	gag Glu	tca Ser	ata Ile	act Thr 15	gcc Ala	ttt Phe	aat Asn	cta Leu	aca Thr 20	att Ile	Gly ggg	wga Xaa	315
Phe P	ca tat ro Tyr 5						_	gg							344
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Met Cys Arg Ala Ala Cys Ile Ile Arg Met Ala Val Arg Ile  -30 -25 -20	0,5
tca ttc ttt ctt tct tac cat gct ctg tct ctc tgc ctt tgt aca tgt  Ser Phe Phe Leu Ser Tyr His Ala Leu Ser Leu Cys Leu Cys Thr Cys  -15 -10 -5	17
gcg ttt gca ttt ctc tcc ctc ctc ggg 2 Ala Phe Ala Phe Leu Ser Leu Leu Gly 1 5	44
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-15	
ctg gma gcc ctc gga ttc ctg amc cag gtg aat ccc arc cca att sma 1 Leu Xaa Ala Leu Gly Phe Leu Xaa Gln Val Asn Pro Xaa Pro Ile Xaa -10 -5 1	.02
ggd ggg tca aaa atg tgt gag twa cac ccc agg ata ctg cag gac atg 1 Gly Gly Ser Lys Met Cys Glu Xaa His Pro Arg Ile Leu Gln Asp Met 5 10 15 20	.50
	.98
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85

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1

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<222> 189
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                                                                       60
cagagettet tegaettate etgeeteece taetttaatt etgttaaagt agttgaacae
                                                                      120
cattettete ataatagtte teectesatt etteagtgat tyeettgtgt ttataggata
                                                                      180
aagtccacnt gttattttgg cagtcagttc aagatccaca aatcagtctt tacccttaca
                                                                      240
tecttattee teactgetgt tetaatatag tetttatace agteaggetg gtetgtteae
                                                                      300
tatteetga atg ttt tte tee att ett ttg tta ttg gea eee eeg eta eee
                                                                      351
         Met Phe Phe Ser Ile Leu Leu Leu Leu Ala Pro Pro Leu Pro
              -15
                                  -10
tct gca gtg tct ttg cta cct ttc ttt ttc tac tgt gtg cag gg
                                                                      395
Ser Ala Val Ser Leu Leu Pro Phe Phe Phe Tyr Cys Val Gln
                        5
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caactctgct gttttgtagg aagccacatg gaggtcattt acggttacta gttatcttag
                                                                       60
tcagcttggg cagccattaa aaaataatac tgtagacgga gtggcccaaa cgagagaaat
                                                                      120
ttatttctta tagttttggc atg gta gat ttc atc ctg agg tct ctt ctc ttg
                                                                      173
                      Met Val Asp Phe Ile Leu Arg Ser Leu Leu
                              -20
                                                   -15
gath tigh agt tigg ofg toa aho too otg cat got, cad acg adought the
                                                                      221
Val Cys Ser Trp Leu Ser Ile Ser Leu His Ala His Thr Thr Ala Phe
                        -5
tgt aca tac agt aag aaa ata cac act gtc atg tca ttt ttt tgt aa
                                                                      268
Cys Thr Tyr Ser Lys Lys Ile His Thr Val Met Ser Phe Phe Cys
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<210> 140

87 <211> 170 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 93..170 <221> sig_peptide <222> 93..140 <223> Von Heijne matrix score 8.10000038146973 seq LLYFLCVSSYVTS/FF ttttgactga tatcaaattc taggtggacc gagattttct ttcagtcttt caaagatatt 60 actitattgc cttctatctt gcatagtttc tg atg aga agt ctg ttg tat ttc 113 Met Arg Ser Leu Leu Tyr Phe -15 tta tgt gtt tct tca tat gta aca tct ttt ttc ttt ttt ttt ttt ttt 161 Leu Cys Val Ser Ser Tyr Val Thr Ser Phe Phe Phe Phe Phe Phe 170 ttt ttt ttt Phe Phe Phe 10 <210> 141 <211> 396 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 192..395 <221> sig peptide <222> 192..236 <223> Von Heijne matrix score 8 seq FISFLCLIALAGT/SS <400> 141 gatteteage ttagttgetg ttggtgtata ggagagetae tgatttgtgt acattaattt 60 tgtatccgga aactttgttg aattatttta tcagttctag gagctttttg gaggagtctt 120 180 tagggttctc taggtataca atcatatcat cagcaaacag tgacaattcg acttcctctt tatggatttg t atg ccc ttt att tct ttc ctt tgt ctg att gct ctg gct 230 Met Pro Phe Ile Ser Phe Leu Cys Leu Ile Ala Leu Ala -10 ggg act tcc agt act atg ttg aga agt gct ctg gct ggg act tcc agt 278 Gly Thr Ser Ser Thr Met Leu Arg Ser Ala Leu Ala Gly Thr Ser Ser act atg tkg arg aga agt ggt gam agt ggg wat cct kgh ctk gty cma 326 Thr Met Xaa Xaa Arg Ser Gly Xaa Ser Gly Xaa Pro Xaa Leu Val Xaa

gtc ctm aga ggg aat gct ttc agc ttt ttc cca ttc agt ctg atg twg

Val Ten Arg Gly Asn Ala Phe Ser Phe Phe Pro Phe Ser Len Met Xaa

35

40

45

gct atg ggt tgt cat aga tgg c

396

Ala Met Gly Cys His Arg Trp

20

<210> 142

<211> 357

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tgttctgcaa taggcggctt agagggaggg gctttttcgc ctatacctac tgtagcttct
                                                                       120
ccacgtatgg accetaaagg ctactgctgc tactacgggg ctagacagtt actgtctcag
                                                                       180
ctctaggatg tgcgttcttc cactagaagc tcttctgagg gaggtaatta aaaaacagtg
                                                                       240
gaatggaaaa acagtgctgt agtcatcctg taatatgctc cttgtcaaca a atg tat
                                                                       297
                                                           Met Tyr
aca ttc ctg cta ggt gcc ata ttc att gct tta agc tca agt cgc atc
                                                                       345
Thr Phe Leu Leu Gly Ala Ile Phe Ile Ala Leu Ser Ser Ser Arg Ile
                 -10
tta cta gtg aag
                                                                       357
Leu Leu Val Lys
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<223> Von Heijne matrix
       score 7.90000009536743
       seq LVCVCVCVCVCXC/XR
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tgtgtgtgtg tgtgtctgcg tgtgt atg tgt ttg tgt ccc tgc tgg gat gtg
                                                                        52
                             Met Cys Leu Cys Pro Cys Trp Asp Val
                                     -40
                                                                       100
ttt act gtg ttt gtg tgt gtc tct gtg tgt gtg tct gtg tct gtc cct
Phe Thr Val Phe Val Cys Val Ser Val Cys Val Ser Val Ser Val Pro
                                 -25
gtc ggg atg tat tta gtg tgt gtg tgt gtg tgt gtg tgt gtg tgt stc
                                                                       148
Val Gly Met Tyr Leu Val Cys Val Cys Val Cys Val Cys Val Cys Xaa
         -15
                             -10
                                                                       159
tge gyg egt gg
Cys Xaa Arg
. , 1
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 <212> DNA
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<222> 282..383
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      seg LFSLLMLTQSPLA/GQ
<221> misc feature
<222> 132,149
<223> n=a, g, c or t
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                                                                       60
gtggeececa ttggeeatgg etgeagagee gatgteeegg ecaateeagg egggateeee
                                                                      120
ttgaagcmgg knsmwhbtcy kragscwknc cmabtctccg ggggcaastc ttttcccttc
                                                                      180
cctgtgaccc kcttcggaca gttgaccatc tcaacaccta gtggttaaaa agaagagcat
                                                                      240
ggacggcctg gggcctgcac tggctgtgct gggagtttgt c atg ttg ata gct aag
                                                                      296
                                               Met Leu Ile Ala Lys
                                                               -30
cag gcc cag ccc caa ggc ctc act gcc atc tgc ttc cct ctc aca cct
                                                                      344
Gln Ala Gln Pro Gln Gly Leu Thr Ala Ile Cys Phe Pro Leu Thr Pro
                -25
                                     -20
ctc ttc tcc ctc ctc atg ctc act cag agc ccc ctt gca ggt cag gaa
                                                                      392
Leu Phe Ser Leu Leu Met Leu Thr Gln Ser Pro Leu Ala Gly Gln Glu
            -10
                                -5
gga aga gaa ggg aaa gaa cgg tac ttg ttg gtg att ca
                                                                      433
Gly Arg Glu Gly Gly Lys Glu Arg Tyr Leu Leu Val Ile
                        10
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                                                                       50
                Met Leu Arg Thr Trp Ser Ser Leu Pro Trp Thr Arg
                    -25
                                         -20
ttt egg gtt tge ttg etc tet etc tet etc ttt etc tgg get aat egt
                                                                       98
Phe Arg Val Cys Leu Leu Ser Leu Ser Leu Phe Leu Trp Ala Asn Arg
                -10
                                     -5
                                                                      146
tta gag gad agt ege ted tge daa det aat dee atg age etg adt acc
Leu Glu Asp Ser Arg Ser Cys Gln Pro Asn Pro Met Ser Leu Thr Thr
                            Τ0
ttq ccq qqc cac agg ctc aaa gaa gca gtg tgg ctg cca gca ccc tca
                                                                      194
Leu Pro Gly His Arg Leu Lys Glu Ala Val Trp Leu Pro Ala Pro Ser
                        25
ctt ggg
                                                                      200
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Leu Gly

90

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<222> 80..166
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                                                                       60
eggeacaatt taattaetg atg gee eet tte eta ega eag gtg gat rtg tgg
                                                                      112
                    Met Ala Pro Phe Leu Arg Gln Val Asp Xaa Trp
                                      -25
gga gca cag gcc ggt ctg gtg gtb gsm tgg tta cta cca tgs caa tgc
                                                                      160
Gly Ala Gln Ala Gly Leu Val Val Xaa Trp Leu Leu Pro Xaa Gln Cys
            -15
                                -10
age tgt gaa ega tea gag eaa tat etg age ace tgt ete eea eag eae
                                                                      208
Ser Cys Glu Arg Ser Glu Gln Tyr Leu Ser Thr Cys Leu Pro Gln His
                                             10
tca agc atc aag cag tcg tgc atc aag cat cca gca ggc ccg atc ccc
                                                                      256
Ser Ser Ile Lys Gln Ser Cys Ile Lys His Pro Ala Gly Pro Ile Pro
                    20
                                        25
gca ggc cac cta cag gga aag gcc aca gct gcg ccc ctg gg
                                                                      297
Ala Gly His Leu Gln Gly Lys Ala Thr Ala Ala Pro Leu
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                                                                       60
tgcacagaga gaactcacc atg gag ttt ggg ctg aag tgg ctt ttt ctt gtg
                                                                      112
                     Met Glu Phe Gly Leu Lys Trp Leu Phe Leu Val
gca att tta aaa ggt gtc cgg tgt gaa gtg aag ctg gtg qag tct ggg
                                                                      160
Ala Ile Fen Lys Gly Val Arg Cys Glu Val Lys Leu Val Glu Ser Gly
            -5
gga ggc ctg gtg cag ccg ggg ggg tcc ctg aga ctc tcc tgt gta gga.
                                                                      208
Gly Gly Leu Val Gln Pro Gly Gly Ser Leu Arg Leu Ser Cys Val Gly
                        15
tot gga tto gto tto gat aaa tat ggo ata agt tgg gtg cgc cag gca
                                                                      256
Ser Gly Phe Val Phe Asp Lys Tyr Gly Ile Ser Trp Val Arg Gln Ala
25
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<222> 56..115

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seq LLLFPLSLLFTLG/FL

<221> sig_peptide

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93 <222> 69..122 <223> Von Heijne matrix score 7.80000019073486 seq LCVLSLLVSFKSA/CL <400> 151 cacattttct acttaaaagc aamgttacaa agcctgtgga attgctctga cttaqaaaga 60 acttgate atg ctt ttg gag tet eta tgt gtt ete tet etg ttg gtt agt 110 Met Leu Leu Glu Ser Leu Cys Val Leu Ser Leu Leu Val Ser ttt aaa tca gcc tgc ctc aca agg gag cct gca ttt gat tcc caa gcc 158 Phe Lys Ser Ala Cys Leu Thr Arg Glu Pro Ala Phe Asp Ser Gln Ala cgc ccg gg 166 Arg Pro <210> 152 <211> 382 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 99..380 <221> sig_peptide <222> 99..236 <223> Von Heijne matrix score 7.80000019073486 seq LLYLSFAALGVVA/LR <400> 152 ttttacacac acacatacat acacacacac agctaattga gttttaaagt aatattcttg 60 ctaatcccta ctgaattgta gcttggtgtt gtttctga atg gtt ttt gga tat tgg 116 Met Val Phe Gly Tyr Trp -45 aag cag ccg ctg att acc ctt gca aag aaa tct gta aaa tgt gca cgt 164 Lys Gln Pro Leu Ile Thr Leu Ala Lys Lys Ser Val Lys Cys Ala Arg -35 -30 gaa tgt ctg aga tgc tct ctc agg cct cta gtc ctt ctg tat ctt tcc 212 Glu Cys Leu Arg Cys Ser Leu Arg Pro Leu Val Leu Leu Tyr Leu Ser -20 -15 ttt gca gcc ctg ggt gta gta gca ctc agg agt gtt gaa tca ccc ctg 260 Phe Ala Ala Leu Gly Val Val Ala Leu Arg Ser Val Glu Ser Pro Leu gcc gag acc cac tcc tgc tgg ctc agc ctg ggc atg tgt gtg ctc cag 308 Ala Glu Thr His Ser Cys Trp Leu Ser Leu Gly Met Cys Val Leu Gln 15 tgt gaa cag cag tgg gtt cca acc cca gtc tcc ttt ctc tgt ggc ctc 356 Cys Glu Gln Gln Trp Val Pro Thr Pro Val Ser Phe Leu Cys Gly Leu 30 35 40 tet gge tee age ace ate ate gtt ag 382 Ser Gly Ser Ser Thr Ile Ile Val 45 <210> 153 <211> 208 <212> DNA <213> Homo sapiens

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PCT/IB99/00712 WO 99/53051 94

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	ga tgc tgg att ttg tca aat cct ttt tct gca tgt att gag atg ly Cys Trp Ile Leu Ser Asn Pro Phe Ser Ala Cys Ile Glu Met -25 -20 -15	101
	tg tta ttt ttg ttt tta att ctg ttt ata tgg cac att cgg g eu Leu Phe Leu Phe Leu Phe Ile Trp His Ile Arg -10 -5 1	147
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atg go Met Al	la Glu Lys Ala Gly Ser Thr Phe Ser His Leu Leu Val Pro Ile -20 -15 -10	60 108 141
Leu Le	eu Leu Ile Gly Trp Ile Val Gly Cys Thr	
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<400> ctcaa	157 gaagc c atg gcg gaa tcc agg ggc cgt ctg tac ctt tgg atg tgc Met Ala Glu Ser Arg Gly Arg Leu Tyr Leu Trp Met Cys	. 50
ttg go Leu A	-15 -10 get get geg etg gea tet tte etg atg gga ttt atg gtg gge tgg la Ala Ala Leu Ala Ser Phe Leu Met Gly Phe Met Val Gly Trp	98

5

Ala Met Tyr Tyr

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96
ttt att aag cct ctg gg
                                                                       115
Phe Ile Lys Pro Leu
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                                                                       104
Leu Thr Ser Leu Pro Phe Leu Leu Pro Thr Ile Ser Phe Leu Leu Leu
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                                         -15
ttg tat ttt ttt cma att gct gtc acc cat ccg tca gtt ctc atc aac
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Leu Tyr Phe Phe Xaa Ile Ala Val Thr His Pro Ser Val Leu Ile Asn
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                                                                       120
atg aga aaa gad gtg agg tite off titg tite off, acc tigh ggd off off.
                                                                       168: . . .
Met Arg Lys Asp Val Arg Phe Leu Leu Phe Phe Thr Cys Gly Leu Pro
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                                         -10
gcc cta cac ggg gac tct agg gtg gaa tgt agc aaa gcc cat cca cca
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Ala Leu His Gly Asp Ser Arg Val Glu Cys Ser Lys Ala His Pro Pro
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gcc atg tac tac cc
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gcc caa ggc ctg acc cag act ccg acc gaa atg cag cgg gtc agt tta Ala Gln Gly Leu Thr Gln Thr Pro Thr Glu Met Gln Arg Val Ser Leu

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                                                                      100
Leu Ile Ser Glu Leu Leu Leu Arg Ser Val Thr Ser His Asn Thr
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Met Met Arg Ala Leu Ser Ser Gln Met Leu Ser Gln Ser Phe Pro Arg
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Pro Ser Phe Gly Phe Ile Ser Lys Ile His Pro Ser His Pro Pro
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                                          -50
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                                                                      102
Met Phe Ile Val Val Met Val Gln Ile Cys Gly Arg Asn Gly Lys Arg
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Ser Asn Arg Thr Leu Arg Glu Glu Val Leu Arg Asn Leu Arg Ser Val
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-15

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-30	-25	-20	

PCT/IB99/00712 WO 99/53051 101 car ctc ctt tta ctg aca tcc cat ttt cta ggc gag tcc ctt ggt gga 151 Gln Leu Leu Leu Thr Ser His Phe Leu Gly Glu Ser Leu Gly Gly -10 ggc aca ctg ctt gtc cca ctc ctc ccc cca ggg 184 Gly Thr Leu Leu Val Pro Leu Leu Pro Pro Gly <210> 168 <211> 218 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 97..216 <221> sig_peptide <222> 97..177 <223> Von Heijne matrix score 7.40000009536743 seq ILLLTICAAGIXG/TR <400> 168 cettecetee gegeacagge tgeeggetea cegettgeta atggeageeg gggteteeet 60 gggacagcaa gacctccgct caggcccctc tttcga atg ckc cam gcm ctc ctg 114 Met Xaa Xaa Ala Leu Leu cga tot aga atq att cag ggc agg atc ctg ctc ctg acc atc tgc gct 162 Arg Ser Arg Met Ile Gln Gly Arg Ile Leu Leu Thr Ile Cys Ala -15 -10 gcc ggc att rgt ggg act cgt cag ttt ggc tat aac ctc tct atc atc 210 Ala Gly Ile Xaa Gly Thr Arg Gln Phe Gly Tyr Asn Leu Ser Ile Ile 218 aat gac cc Asn Asp <210> 169 <211> 480 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 317..478 <221> sig_peptide <222> 317..457

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Met Met Leu Asp Phe Ala Leu Ser Pro

-35
-30

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Arg Leu Glu Arg Ser Gly Leu Ile Met Ala Cys Cys Thr Leu Asp Leu

-25

-20

-15

390

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Gln Val Tyr Tyr Val Ser Gln Leu Ile 25 30

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tct tcc tcc cca tca ggg gca g Ser Ser Ser Pro Ser Gly Ala V			
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aat gga att ttc ttg ctc ttg atc tct gtc tta aca gtg att tgg ttt Asn Gly Ile Phe Leu Leu Ile Ser Val Leu Thr Val Ile Trp Phe -15 -10 -5	227
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gct tt Ala Le	tg tt eu Ph	c ie	ttt Phe	ttg Leu -10	ctt Leu	agg Arg	att Ile	gct Ala	ttg Leu -5	gct Ala	agt Ser	tgg Trp	gct Ala	ctc Leu 1	ttt Phe	162
tgg at Trp I	tc ca le Hi 5	t .s	atg Met	aat Asn	ttt Phe	aga Arg	aga Arg 10	gct Ala	ttt Phe	ttc Phe	cac His	tta Leu 15	cgg Arg	tgg Trp	ttt Phe	210
gat at Asp II	le As	it sn	agc Ser	act Thr	gaa Glu	tct Ser 25	gta Val	aat Asn	tgc Cys	ttt Phe	30 GJA aaa	cag Gln	tat Tyr	ggc Gly	cta Leu	258
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109

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Gly Asp Lys Leu Gly Asp Lys Tyr Ala Cys Trp Tyr Gln Gln Lys Pro	
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Gly Gln Ser Pro Val Leu Val Ile Tyr Gln Asp Ser Lys Arg Pro Ser	
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Gly Ile Pro Glu Arg Phe Ser Gly Ser Asn Ser Gly Asn Thr Ala Thr	
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Leu Thr Ile Ser Gly Thr Gln Ala Met Asp Glu Ala Asp Tyr Tyr Cys	
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                                                                      105
Ile Leu Tyr Leu Phe Phe Leu Lys Trp Ser His Pro Gly Trp Ser
    -15
                        -10
                                             -5
gea acg ncg tgg tet tgg cac act gea acc tee gee tee etg att caa
                                                                      153
Ala Thr Xaa Trp Ser Trp His Thr Ala Thr Ser Ala Ser Leu Ile Gln
                                     10
gtg att ctc ccg cct tgg g
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Val Ile Leu Pro Pro Trp
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tgt ttt ctg caa atg gac aat ttg act cct ctt ttc cta tct gga tgc
                                                                      103
Cys Phe Leu Gln Met Asp Asn Leu Thr Pro Leu Phe Leu Ser Gly Cys
                                                     -10
            -20
                                -15
                                                                      150 -
the tha fit of for cwt tgo wtg att tab ttg got agg atr ttg gg-
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 agageteetg teeggtgtge cageageeeg gaetggeggt gagegegagg gaggetaekg
                                                                        120
 agaagcccgg cgacggagga acgcaggtct gctgccaggg attgaggaga ctgaagaacg
                                                                        180
 ctgaagacag gctg atg ggc tca gct ggt agg ctc cac tat ctc gsc atg
                                                                        230
                 Met Gly Ser Ala Gly Arg Leu His Tyr Leu Xaa Met
                                      -35
 act gct gaa aat ccc act cct gga gac ctg gct ccg kcc ccc ctc atc
                                                                        278
 Thr Ala Glu Asn Pro Thr Pro Gly Asp Leu Ala Pro Xaa Pro Leu Ile
             -25 ·
                                  -20
 act tgc aaa ctc tgc ctg tgt gag cag tct crt gga caa gat gac cac
                                                                        326
 Thr Cys Lys Leu Cys Leu Cys Glu Gln Ser Xaa Gly Gln Asp Asp His
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 act cca gga atg c
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                                                                       113
                                         Met Asn His Leu Pro Pro
 aso cat tat agg mgc cat gtg ttc aca tgt cat gtg gac cag tal tta
                                                                       161
 Asn His Tyr Arg Xaa His Val Phe Thr Cys His Val Asp Gln Tyr Leu
             -40.
                                - 35
 act gtg gaa acc gcg ggt ggc atg gag aag gag gca gtg tcc gtg act
                                                                       209
 Thr Val Glu Thr Ala Gly Gly Met Glu Lys Glu Ala Val Ser Val Thr
         -25
                                                  -15
                             -20
 gtg ctg ctc tcc gca gcc ccc tgc ctg ctg tcc tgt ttc ctc ggc tcc
                                                                       257
 Val Leu Leu Ser Ala Ala Pro Cys Leu Leu Ser Cys Phe Leu Gly Ser
                         -5
                                                                       305
 tog gtg tot gga otg gcg tto tgg gtt too cag cag aaa act aaa ggg
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Ser Va	l Ser	Gly	Leu 10	Ala	Phe	Trp	Val	Ser 15	Gln	Gln	Lys	Thr	Lys 20	Gly	
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gcg ag	_														359
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	138	257 eijn 7	e ma		/CT										
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aca ca Thr Hi															218
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	147	272 eijn 7	e ma		/нр										
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ttc ag Phe Ar															221
cat at His Il		aaa Lys					Ser								269

114	
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cat cgt ccc agg agc agc acc agc tac agg aac ctg ccg cat ctg ttt  His Arg Pro Arg Ser Ser Thr Ser Tyr Arg Asn Leu Pro His Leu Phe  -25  -20 -15	164
ctg ttt ttc ctc ttc gtg gga ccc ttc agc tgc ctc ggg agt tac agc Leu Phe Phe Leu Phe Val Gly Pro Phe Ser Cys Leu Gly Ser Tyr Ser	212
cgg Arg	215
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ggg ghc agc cgc cta ctg ctg ctg ctg ctg ctg ccg ctg cct Gly Xaa Ser Arg Leu Leu Leu Leu Leu Leu Leu Xaa Leu Pro Leu Pro -15 -10 -5	158
ccg ccg gkv ctg cga acc cgg gdy ttt tca wgc acc aca ctc acc gcm Pro Pro Xaa Leu Arg Thr Arg Xaa Phe Ser Xaa Thr Thr Leu Thr Ala	206

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atg aat ttg ggg gga cat tca gat cat agc act ttt ctt ttc ttt ctt
                                                                      107
Met Asn Leu Gly Gly His Ser Asp His Ser Thr Phe Leu Phe Phe Leu
                            -15
ttt ttt tct gtt ttt tgt ttt ttt t
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Phe Phe Ser Val Phe Cys Phe Phe
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                                                       -20
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gcg atc ttc gcc gtt acc ttc ttg ctg gcg ttg gtg gga gcc gtg ctc
Ala Ile Phe Ala Val Thr Phe Leu Leu Ala Leu Val Gly Ala Val Leu
                            -10
                                                                      153
tac etc tat eeg get tee aga caa get gea gga att eea ggg att act
Tyr Leu Tyr Pro Ala Ser Arg Gln Ala Ala Gly Ile Pro Gly Ile Thr
                                        10
cca act gaa gaa aaa gat ggt aat ctt cca gat att gtg aat agt gga
                                                                      201
Pro Thr Glu Glu Lys Asp Gly Asn Leu Pro Asp Ile Val Asn Ser Gly
                20
                                    25
agt ttg cat gag tbc ctg gtt aat ttg cat gag aga tat ggg cct gtg
                                                                      249
Ser Leu His Glu Xaa Leu Val Asn Leu His Glu Arg Tyr Gly Pro Val
                                40
                                                                      297
gto too tto tgg tit ggc agg cgc oto gtg gtt agt ttg ggc act gtt
Val Ser Phe Trp Phe Gly Arg Arg Leu Val Val Ser Leu Gly Thr Val
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                                                                      112
                               Met Tyr Thr Asn Lys Tyr Thr Leu
                                   -45
ata tat aac ata cta ata tat aat ata tgt btk drg tat atg tgg ttg
                                                                      160
Ile Tyr Asn Ile Leu Ile Tyr Asn Ile Cys Xaa Xaa Tyr Met Trp Leu
            -35
                                -30
ata etc att tat atg tae eta eat att tge etc ttt tgt tge wet ttt
                                                                      208
Ile Leu Ile Tyr Met Tyr Leu His Ile Cys Leu Phe Cys Cys Xaa Phe
                            -15
                                                 -.10
att tot too tgo aat tot gtg ttt coo tgt gtg att atb ttt ott ctg
                                                                      256
Ile Ser Ser Cys Asn Ser Val Phe Pro Cys Val Ile Xaa Phe Leu Leu
cct gaa gaa ctt ctt twt gtd twt ctd wdw dtg tnt tty wtt gtg aga
                                                                      304
Pro Glu Glu Leu Leu Xaa Val Xaa Leu Xaa Xaa Yaa Phe Xaa Val Arg
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Trp Ser Leu Xaa Xaa Ser Ser Arg Leu Glu Cys
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                          Met Leu Leu Thr His Asn Glu Asp Tyr Met
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Pro Gly Asn Xaa Xaa Xaa Xaa Leu Trp Ser Leu Ile Gln Ala Val
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gct a Ala I																161
ggg g	ly 1															209
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cma g Xaa G																305
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	ata	ca g	tgat	cctc	ca to	caaca	aatt	ato	aaag gga a	jaac itg a	tgta ag a ys A	itgag igg t	ıga a	aaagg	tctct tg	240 293
ttt g Phe V											gtc	acc				341
tgc t Cys C																389
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-35	
ctt ctg gcc tgt aag gtt ttc act gaa aag tct cct acc aaa cat att Leu Leu Ala Cys Lys Val Phe Thr Glu Lys Ser Pro Thr Lys His Ile -30 -25 -20	224
aga gag cac cat tgt atg tta ttt gtt tct ttt ctc ttg ctg ctt tta Arg Glu His His Cys Met Leu Phe Val Ser Phe Leu Leu Leu Leu	272
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Met Thr	- <del></del>
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age tge tge caa ggg gcc tge tge cea tet aca cet cae gag gge act	214
Ser Cys Cys Gln Gly Ala Cys Cys Pro Ser Thr Pro His Glu Gly Thr -5 1 5	
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                                                          Met Val
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tca tcc ttt acg tgg tgg gca cct gcc tgc tgt gct cca cgt aca tac
                                                                       104
Ser Ser Phe Thr Trp Trp Ala Pro Ala Cys Cys Ala Pro Arg Thr Tyr
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     -10
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Val Val Ser Ala Thr Thr Leu Ser Ala Val Gln Gly His Cys Pro Leu
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att ggg gga ag Ile Gly Gly Ly -15				g gag tgc ctt n Glu Cys Leu	
gcc ctg cct ga Ala Leu Pro G		gtg ttc tg	t agg ggt gg	c tgc aca gcc	
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Val Arg Gln A	gct aac atc la Asn Ile 25	agg atg ca Arg Met Gl -2	n Cys Lys I	to tat gat too le Tyr Asp Ser -15	ctg 286
				gr ctg atg tgt	gct 334

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                                      -35
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                                                     -15
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                                                                       102
Leu Phe Val Leu Val Gly Ser Leu His Leu Phe Leu Ser Val Leu Ala
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                         -10
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Gly Lys Arg

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		ig_pe		le													
	> Vo	on He	eijne 6.69	9999	rix 98092 XXXP/		ŀ										
	se	eq FC	ALLLI	יעתכי	\AAP/	AA											
	)> 2]		ana.	tcca	aa o	ato	aca	gga	tcc	agg	caa	agg	aat	ctc	caa		51
	.gcgt		ague		iu g			Gly									71
-	_	_		_			-	gcc Ala		_	_			_			99
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aca	tcg	ccg	ı ttt	cga	gta	saa	ata	cag	ctt	caa	999	10 gcc	ġca	cct	ggt		195
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30		_	_	_	35		•		_	40				-	45		
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261	цуз	FILE	Arg	50	ASII	Jei	Der	Ser	55	116	·	цуз	цуз	60	nys .	•	
		_		_		_		agc Ser 70									327
-210	)> 2:	17												•			
	.> 35																
	> Di > Ho	NA omo s	sapie	ens				•									
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								ctg									159
Lys -5	GTÀ	val	HlS	cys	Asp 1	vaı	GID	ьeu	vaı 5	GIU	ser	стА	дтÀ	10	Leu	•	•
			Gly					ctc Leu					Ser			;	207
acc	ctc	agt	15 aac	gac	tgg	atg	cac	20 tgg	gtc	cgc	caa	gcc	25 cca	999	aag	:	255

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-20

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gtc ttc aag gac tgg att cga gac cag ctc aac ct Val Phe Lys Asp Trp Ile Arg Asp Gln Leu Asn Le 5 10 15	cc ttc atc aac aac 247 eu Phe Ile Asn Asn 20
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score 6.5

seq WLFFLMLSLCTPP/DR

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ctg ggt gtg aag ccg agc acg cat tgg cta ttc ttc ctg atg ctc tcc Leu Gly Val Lys Pro Ser Thr His Trp Leu Phe Phe Leu Met Leu Ser	148
ctt tgc acc cct cct gac aga ccc tgg tgt gtg ttg ttc ccc ccg ctg Leu Cys Thr Pro Pro Asp Arg Pro Trp Cys Val Leu Phe Pro Pro Leu -5 1 5 10	196 198
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ggg aag aga ggc cag gca tgg cgg ctc atg cct gtw gtc cca gca gtt	160

Gly Lys Arg Gly Gln Ala Trp Arg Leu Met Pro Val Val Pro Ala Val tgg gag cct gag gca ggt gga ttg ctt cag ctc ggg ggt tct agg g 206 Trp Glu Pro Glu Ala Gly Gly Leu Leu Gln Leu Gly Gly Ser Arg 25 20 <210> 228 <211> 480 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 216..479 <221> sig_peptide <222> 216..326 <223> Von Heijne matrix score 6.5 seq LLVFFLIVRTLSC/RS <400> 228 gcatcccck ktagctcaga gaagtttgtt rdgaccgatc ttctgaagcc tacttctgtc 60 aactcatcaa agtcattctc catccagctt tgttccatta tgggtgagga gctacgatcc 120 tttggaggag aagaggcact ctgattttta gaattttcag cttttctgct ctggtttcgc 180 cccatctttg tggttttatc taccttcggt ctttg atg atg gtg acc tac aga 233 Met Met Val Thr Tyr Arg -35 tgg ggt ttt ggt gtg gat gtc mtt ttt gtt gct gtt gat gct att cct 281 Trp Gly Phe Gly Val Asp Val Xaa Phe Val Ala Val Asp Ala Ile Pro -25 ttc tgt ttg tta gtt ttc ttt cta ata gtc agg acc ctc agc tgc agg 329 Phe Cys Leu Leu Val Phe Phe Leu Ile Val Arg Thr Leu Ser Cys Arg -15 -10 -5 tet gtt gga gta tge tgg agg tee aet eea gae eet gtt tge eta ggt 377 Ser Val Gly Val Cys Trp Arg Ser Thr Pro Asp Pro Val Cys Leu Gly 10 atc acc agc aga ggc tgc aga aca gaa ata ttg cag aac agc aaa tgt 425 Ile Thr Ser Arg Gly Cys Arg Thr Glu Ile Leu Gln Asn Ser Lys Cys 25 tgc tcc ctg atc ctt cct ctg gaa gct tcg tct caa agg ggc act gaa 473 Cys Ser Leu Ile Leu Pro Leu Glu Ala Ser Ser Gln Arg Gly Thr Glu 35 40 45 480 tgt atg a Cys Met 50 <210> 229 <211> 144 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 43..144 <221> sig_peptide <222> 43..99 <223> Von Heijne matrix

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score 6.5

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132

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gtaaagagag aagtaatgga aggcctgtct g atg ttg ctt ctt ttg caa cta

Met Leu Leu Leu Gln Leu

-10

aac tta aaa aca ctc tca tcc agt acc ata gca ttg aag aag ata agt

Asn Leu Lys Thr Leu Ser Ser Ser Thr Ile Ala Leu Lys Lys Ile Ser

-5

1

ggc gag ttg cta aga aaa cga aag agg g

Gly Glu Leu Leu Arg Lys Arg Lys Arg

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cwtggtargg tcaytaagaa taactckaat cawgatgtta aaaggctttc ctttacattc
                                                                       180
acaaaacaat ttrsttccta gaagtagttt attcttgcct gtggtcattt ttgctccttt
                                                                       240
ataatactac atctaaatca atttgttaaa tatagtagag aaatgaaata aatttcttcc
                                                                       300
agttaaacca ctgcacttaa agagtagaaa ccctctct atg tca ctc ttt gtt ttg
                                                                       356
                                           Met Ser Leu Phe Val Leu
                                           -15
ttg atc ata act caa ctg ctg tat ggt ggg ata ctc t
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Leu Ile Ile Thr Gln Leu Leu Tyr Gly Gly Ile Leu
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                                                                       60
gttttggtct atagatgtct cattgcttct gtgctatttg ttggaaaagc tgttcttcca
                                                                      120
atg aat tgc ttt tgc aat ttt gtc aaa acc agt gag gca tat atg att
                                                                      168
Met Asn Cys Phe Cys Asn Phe Val Lys Thr Ser Glu Ala Tyr Met Ile
            -25
                                 -20
ctg ttt cta ggt gtt cta ctc tct gca agt gat tta tgt gtc tat ccc
                                                                      216
Leu Phe Leu Gly Val Leu Leu Ser Ala Ser Asp Leu Cys Val Tyr Pro
        -10
                             -5
atc ggg
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Ile Gly
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tot gta atc cta gca ctt tgg gag gcc gag gcg ggc gga tcg cct gag
                                                                       104
Ser Val Ile Leu Ala Leu Trp Glu Ala Glu Ala Gly Gly Ser Pro Glu
             -10
atc ggg agt tcg gga ccg gcc gca cca aca tgg aga agc ccc gtc cag
                                                                       152
Ile Gly Ser Ser Gly Pro Ala Ala Pro Thr Trp Arg Ser Pro Val Gln
gg
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                                                                      120
ctitgcctat tctggacatt ttgtataaaa gggtttgtgg aggatgtggt cttttgtgac
                                                                      180
tggcttcttg aacttggcat agtgttttca aggttcaacc atgttgtagc acgtacgttc
                                                                      240
ctttttatgg ccaa atg tac gga gag tcc aca ttg ttt atc cat tca tca
                                                                      290
                Met Tyr Gly Glu Ser Thr Leu Phe Ile His Ser Ser
                                -25
gtt cat ggg cat ttg ggt tgt ctc ctc ttg gct gtt agg agt agt gct
Val His Gly His Leu Gly Cys Leu Leu Leu Ala Val Arg Ser Ser Ala
        -15
                            -10
                                                 ~5
act gtg aac att acg tac chn nkw gtk tgt gtg gac att cak ntt cat
                                                                      386
Thr Val Asn Ile Thr Tyr Xaa Xaa Val Cys Val Asp Ile Xaa Xaa His
ttc cat atg ctt atg tct gga att act ggg tca tat ggc aac tct ctt
                                                                      434
Phe His Met Leu Met Ser Gly Ile Thr Gly Ser Tyr Gly Asn Ser Leu
tca ct
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Ser
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-15

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138 tgg ten gee eec act tee eea tst ace tet tgt eet eec eec aac ace 301 Trp Ser Ala Pro Thr Ser Pro Xaa Thr Ser Cys Pro Pro Asn Thr asc acc aca ccg gyt cc 318 Xaa Thr Thr Pro Xaa 15 <210> 241 <211> 405 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 123..404 <221> sig peptide <222> 123..257 <223> Von Heijne matrix score 6.30000019073486 seq GFVSLLVVHAADA/WV <400> 241 tagctggacc cgtctgggag gtaggtttgt gagcgtgaga gaksgatctg taccgcgggg 60 atccgaagta tgcttatcca ggtgggctgc ctcaagcctc gatcccaccc ccgcgctdvt 120 ag atg gtg tca agg tcc ttg cgt ggg aga agg act tgg gtg aga tgc 167 Met Val Ser Arg Ser Leu Arg Gly Arg Arg Thr Trp Val Arg Cys -45 -40 -35 atg cgg aga ttg ccc cca att ccg gcc tgg agc caa ggg aaa ggg atg 215 Met Arg Arg Leu Pro Pro Ile Pro Ala Trp Ser Gln Gly Lys Gly Met -25 -20 cct gga ttt gtg tct cta ttg gtg gtc cac gct gcg gat gcc tgg gta 263 Pro Gly Phe Val Ser Leu Leu Val Val His Ala Ala Asp Ala Trp Val -10 -5 gcc cag agg ttr tct acg cca tac ttc tca ctg ttt ttg agc ata cct 311 Ala Gln Arg Leu Ser Thr Pro Tyr Phe Ser Leu Phe Leu Ser Ile Pro 10 aga tgt tcc ttt cct agg cgg agt ata gat cgc acg tgt tct agc stc 359 Arg Cys Ser Phe Pro Arg Arg Ser Ile Asp Arg Thr Cys Ser Ser Xaa 20 25 30 tta gac tca gag ggt tcg agc tct ata asc ccc tcc act ccc ttc a 405 Leu Asp Ser Glu Gly Ser Ser Ser Ile Xaa Pro Ser Thr Pro Phe 35 <210> 242 <211> 242 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 129..242 <221> sig_peptide <222> 129..191 -- <223> Von Heijne marrix score 6.30000019073486 seq SLLPCSLISDCCA/SN <400> 242 cttttgtttt gcaatgccct gccccagag gtggagtcta cagaggcagg caggcctcct 60 tgagctgagg tgggctccac ccagttcgag cttcccagct gctttgttta cctactcaag 120

cctgggca atg gtg ggc gcc ctt ccc cca gcc tcg ctt ctg cct tgc agt

139 Met Val Gly Ala Leu Pro Pro Ala Ser Leu Leu Pro Cys Ser -15 -10 -20 ttq atc tca gac tgc tgt gct agc aat gag cga ggc tcc atg ggc gta 218 Leu Ile Ser Asp Cys Cys Ala Ser Asn Glu Arg Gly Ser Met Gly Val -5 gga ccc tct gag cca cgg cgy ggg 242 Gly Pro Ser Glu Pro Arg Arg Gly 10 <210> 243 <211> 363 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 298..363 <221> sig_peptide <222> 298..357 <223> Von Heijne matrix score 6.30000019073486 seq LGSLIASLAPSTG/LG <400> 243 accactctga ggagacgcgt gacagataag aagggctggt gggatcagtc ctggtggtag 60 ctcaggaagc agagcctgga gcatctccac tatggcctgg gctccactac ttctcaccct 120 cetegeteae tgcacaggtt ettgggeeaa etttatgetg acteageege actetgtgte 180 ggagtcgccg gssgaagacg gtaaccatct cctgcacccg cagcagtggc agctttgtca 240 gcaactatgt tcagtggtac cagcggcgcc cggacagtgc ccccaccact gtgatct 297 atg agg atg aca aaa gac cct ctg ggg tct ctg atc gct tct ctg gct 345 Met Arg Met Thr Lys Asp Pro Leu Gly Ser Leu Ile Ala Ser Leu Ala -10 -15 363 cca tcg aca ggt ctt ggg Pro Ser Thr Gly Leu Gly <210> 244 <211> 324 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 153..323 <221> sig_peptide <222> 153..236 <223> Von Heijne matrix score 6.30000019073486 seq FFLFLFFXEXXXX/XX <400> 244 aattgatact gctttagatg tttctgtctc aftttacaaa aatgtaagaa aaaagaaaaa 60 tcaaactata ctgttaccta tttcttgtat attcttaaca gaatgttctg tacacataag 120 tgtatgtgtg traatcotct tgtttaaatg co atg aaa ott cag ott goo ott . 173 Met Lys Leu Gln Phe Ala Phe -25 tgt tat ttt ctt tat tta gat acc ttt ttt ctt ttt ctt ttt ttk 221 Cys Tyr Phe Leu Tyr Leu Asp Thr Phe Phe Leu Phe Leu Phe Phe Xaa -20 -15 -10 gag ama gyc tkg cyc kgt kgc hta ggm agg agt gca gtg gca maa cct 269 Glu Xaa Xaa Xaa Xaa Xaa Xaa Gly Arg Ser Ala Val Ala Xaa Pro

140	
-5 1 5 10	
cag ctc ayt gca gcc tcc acc ttc kgg tty caa gca att tty ctg ccc Gln Leu Xaa Ala Ala Ser Thr Phe Xaa Phe Gln Ala Ile Phe Leu Pro 15 20 25	317
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-65	
aaa ccc ctg agc ggg ctg ctg aat gcg ctg gcc cag gac act ttc cac Lys Pro Leu Ser Gly Leu Leu Asn Ala Leu Ala Gln Asp Thr Phe His -50 -45	101
ggg tac ccc ggc atc aca gag gag ctg cta cgg agc cag cta tat cca Gly Tyr Pro Gly Ile Thr Glu Glu Leu Leu Arg Ser Gln Leu Tyr Pro -40 -35 -30	149
gag gtg cca ccc gag gag ttc cac ccc ttt ctg gca aag atg agg ggg Glu Val Pro Pro Glu Glu Phe His Pro Phe Leu Ala Lys Met Arg Gly -25 -20 -15	197
att ctt aag gta ctg ctc ttt tct gta gtc tcc ggc ttg gag cag aac Ile Leu Lys Val Leu Leu Phe Ser Val Val Ser Gly Leu Glu Gln Asn -10 -5 1	245
ccc ttg gcc gct ggc ttc aga ctc tcc cac ccg gg Pro Leu Ala Ala Gly Phe Arg Leu Ser His Pro 5 10 15	280
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-30 -25 -20 ctg ctg gcg cas tct ctg att ctc tct ccc tct ccg cgt cca gtg ctg	159

141	
Leu Leu Ala Xaa Ser Leu Ile Leu Ser Pro Ser Pro Arg Pro Val Leu -15 -10 -5	
ggc ttt ttc aga caa gtg cat ctc cta acc agg tca cat ttc agc cgc Gly Phe Phe Arg Gln Val His Leu Leu Thr Arg Ser His Phe Ser Arg  1 5 10 15	207
tgg g Trp	211
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tgtattgtta gctgtagtca ccctactgtg ctattgaata ctagagcttg ttccttctgt	180 240
ctaactgt atg att ata ctc att aac caa ctt ctc ttc atc tgt ccc cca  Met Ile Ile Leu Ile Asn Gln Leu Leu Phe Ile Cys Pro Pro -20 -15 -10	290
Pro Pro Pro Ile Ser Ala Ser Ser Asn Tyr His Phe Thr Leu Tyr Leu  -5  1  10	338
cat gac att aac ttt ttt agc His Asp Ile Asn Phe Phe Ser 15	359
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caagaaacag aaacaaaaca gtcacaaaaa agcataaact gttagcattg atccatgatg a atg act gat gta tta ctt caa ttg cta tta aga gtg tgt tct ccc agg	180 229
Met Thr Asp Val Leu Leu Gln Leu Leu Leu Arg Val Cys Ser Pro Arg -15 -10 -5 1	223.
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tctgcataag accagttatc ccagaaccgt ttgttgaata ggaagttctt ttctcattgc
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                                                                      292
                            Met Gly Leu Phe Leu Cys Cys Ser Leu
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ctg ata ttc tgt ctg gtt gtt cta atc ata act gaa ctg ggc tat ggg
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                                                                      120
                                                                      180
ggaaaggccc tcctccagaa aatgctagaa aacctgagtg gggagctggg gagggagtag
tggactetge tteattgtce ceagtetgea caccecetee eccaceacee caetgeattt
                                                                      240
                                                                      296
cccagctcag ccaaactttc tgannaagac gggcagagnn ctgctgggag atg gga
                                                        Met Gly
tec tgg_gcc_ctg act tgg ctc cat cca gca qaq gct ggg acc agg gtg- .-
                                                                      344
Ser I'rp Ala Leu Thr Trp Leu His Pro Ala Glu Ala Gly Thr Arg Var
        -10
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cct tic tgc agc tgg gaa aaa tca gat ggg cgc tct ta
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Ile Tyr Ile Xaa Leu Xaa Asp Leu Tyr Asp Phe Phe Leu Leu Gly Thr

PCT/IB99/00712

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328

-15

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ggatyyatdg ttttgggctt gaaggggttt cacggctttt tctataacaa cg atg gca
                                                                       238
                                                            Met Ala
                                                                -25
tot toa atg ctg waa too tto cag act tto atg atg ttg act cta ttg
                                                                       286
Ser Ser Met Leu Xaa Ser Phe Gln Thr Phe Met Met Leu Thr Leu Leu
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                                                                       120
tragestreet etreataact gtggraggga cacttazece ttoretgget gtgagaagtt
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attctctgag ggctggtgag caga atg gga aga tct aag agg cag ctc ctt
                                                                      231
                           Met Gly Arg Ser Lys Arg Gln Leu Leu
                           -20
tee ttg cet ggt tee ttt ate eet ggg aat tge agg eea agg att etg
                                                                      279
Ser Leu Pro Gly Ser Phe Ile Pro Gly Asn Cys Arg Pro Arg Ile Leu
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10

236

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ttc acc cag tgc tgc ctt att gga ctc ctt gtg cct ctc ctt ggc tgg  Phe Thr Gln Cys Cys Leu Ile Gly Leu Leu Val Pro Leu Leu Gly Trp  -15  -10  -5
gga aat cag aat aca cag tgg tat ccc act tct aag atg cct gat ggg Gly Asn Gln Asn Thr Gln Trp Tyr Pro Thr Ser Lys Met Pro Asp Gly 1 5 10 15
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tat ttc ttg gtg ctt ctt cgt gtt tta tac act tta caa tgg ggt ggg Tyr Phe Leu Val Leu Leu Arg Val Leu Tyr Thr Leu Gln Trp Gly Gly -10 -5 1 5
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-10 -5 aag caa cct aag tgt tca tca aca aac aaa tgg aca aag aaa atg tgg Lys Gln Pro Lys Cys Ser Ser Thr Asn Lys Trp Thr Lys Lys Met Trp 1 5 10 15	219
tac ata tac aca atg gag tac tat tca gcc ata aaa aaa gat gat atc Tyr Ile Tyr Thr Met Glu Tyr Tyr Ser Ala Ile Lys Lys Asp Asp Ile 20 25 30	267
ctg tca ttt gca aca ata tgg atg gaa ctg gag agc att aca tta agt Leu Ser Phe Ala Thr Ile Trp Met Glu Leu Glu Ser Ile Thr Leu Ser 35 40 45	315
gaa ata agt ggg sca cca aaa gac aaa ctt ctc atg ttc tca ctc att Glu Ile Ser Gly Xaa Pro Lys Asp Lys Leu Leu Met Phe Ser Leu Ile 50 55 60	363
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gcc cac ctg agc atc ctc cag agc ctc gtg cca gct gct ggt gca gyc Ala His Leu Ser Ile Leu Gln Ser Leu Val Pro Ala Ala Gly Ala Xaa -10 -5 1	277
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Met	
ggt tgt ctt ctg gcc tca gag tat ccc tta tca gaa cct tgg gcc cct	167
Gly Cys Leu Leu Ala Ser Glu Tyr Pro Leu Ser Glu Pro Trp Ala Pro	
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Gly Pro Phe Thr Gln Tyr Leu Val Asp His His His Thr Leu Leu Cys	
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aat ggg tat tgg ctt gcc tgg ctg att cat gtg gga gag tcc ttg tat	263
Asn Gly Tyr Trp Leu Ala Trp Leu Ile His Val Gly Glu Ser Leu Tyr	
-45 -40 -35 -30	
gcc ata gta ttg tgc aag cat aaa ggc atc aca agt ggt cgg gct cag	311
Ala Ile Val Leu Cys Lys His Lys Gly Ile Thr Ser Gly Arg Ala Gln	311
-25 -20 -15	
	250
cta ctc tgg ttc cta cag act ttc ttc ttt ggg ata gcg tct ctc asc	359
Leu Leu Trp Phe Leu Gln Thr Phe Phe Phe Gly Ile Ala Ser Leu Xaa	
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Ile Leu Ile	
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aaccctcctt gcatccctgg aatgattcct acttcattat agtgtataat ctttttg	177
atg tgc tgt tgg att tgg gtt gct agt att ttg ttg aga att ttt gca	225
Met Cys Cys Trp Ile Trp Val Ala Ser Ile Leu Leu Arg Ile Phe Ala	
-15 -10 -5	
tot gtg tta atc agg gat att tac ctg tgg ttt tot ttt ttt ttt t	274
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                                                           Met Ile
tcc tca cat tta tat aac ttc agt ctc ctg ttc ttt kta ctc tgg ctg
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Ser Ser His Leu Tyr Asn Phe Ser Leu Leu Phe Phe Xaa Leu Trp Leu
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                         -15
agg tac aag gaa tca gga aga gag ggc aac tgt gag gaa gga gca ttc
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Arg Tyr Lys Glu Ser Gly Arg Glu Gly Asn Cys Glu Glu Gly Ala Phe
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                                                                      109
  Met Gly Trp Gln Arg Leu Leu Leu Leu Pro Arg Pro Pro Ala Ser Thr
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                              -10
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Gly Ala Ser Asn Ala Thr Arg Xaa Pro Lys Xaa Leu Tyr Arg Xaa Tyr
aac cac ggt gtg ctg aag ata acc atc tyt aaa tec tgc cag Reasdot
                                                                      205
Asn His Gly Val Leu Lys Ile Thr Ile Cys Lys Ser Cys Gln Lys Pro
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gta gac aaa tat atc gag tat gat cct gtt atc atc ttg awk aat gct
Val Asp Lys Tyr Ile Glu Tyr Asp Pro Val Ile Ile Leu Xaa Asn Ala
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Ile Leu Cys Lys Ala Xaa Ala Tyr Arg His Ile Leu Phe Asn Thr Gln
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Ser Gly Leu Leu Cys Asn Lys Ser Phe

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cta ttc aag ttc tta gcc cac ttt tta atc ggg tta aca gtt tgt ttt Leu Phe Lys Phe Leu Ala His Phe Leu Ile Gly Leu Thr Val Cys Phe -15 -10 -5	97
ggt gag ggr wgg cta atg agt tat agg agt tct tat tta tta ctt aaa Gly Glu Gly Xaa Leu Met Ser Tyr Arg Ser Ser Tyr Leu Leu Leu Lys 1 5 10 15	145
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c atg tgg tgg ggg aga tgc ttc atc cgg gtc ttg cat ttg ttc cct ctg Met Trp Trp Gly Arg Cys Phe Ile Arg Val Leu His Leu Phe Pro Leu	180 229
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-25 -20 teg tet ttg tte atg gee ett eea eea gtg etg age tea eat gge age	250
Ser Ser Leu Phe Met Ala Leu Pro Pro Val Leu Ser Ser His Gly Ser	278
-15 -10 -5 1	•
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Arg Asn Leu Arg Ile Trp Gly Ser Pro Phe Gly Gly Ala Leu Thr Lys 5 10 15	
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Trp Leu Cys Leu Pro Cys Ser Leu Cys Val Ser Gln Leu Leu Pro Ser -10 -5 1	_

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tgg gca ggc cta tta tcc cta ctt ggc ccg ctc wgt ccg cct atg agg Trp Ala Gly Leu Leu Ser Leu Leu Gly Pro Leu Xaa Pro Pro Met Arg -15 -10 -5	162
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gct gat tta tta ccc agc tca tcc ttt gct aat ccc aag ctg agt ggg Ala Asp Leu Leu Pro Ser Ser Ser Phe Ala Asn Pro Lys Leu Ser Gly -10 -5 1 5	220
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215

220

774

cag at

Gln

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-15 ttt ctg gac aca	gaa gta t	-10 tt qtq aca		age cca	-5 aca cct	tcc 216
Phe Leu Asp Thr						
1	5 toa toa o			10		251
ccc act ggt gtt Pro Thr Gly Val						251
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gccaggag atg aca gca ctg ggg ttt gtt ctg tta gct cca cgt ggc tgg  Met Thr Ala Leu Gly Phe Val Leu Leu Ala Pro Arg Gly Trp  -10  -5	170
-15 -10 -5 ggg agc ctc aca gtc atg gtg gaa ggc aag gaa gag caa gtc acg tct	
Gly Ser Leu Thr Val Met Val Glu Gly Lys Glu Glu Gln Val Thr Ser	218
1 5 10 15	
tac acg gat ggc agc agg caa aga gac agc aat ttt	254
Tyr Thr Asp Gly Ser Arg Gln Arg Asp Ser Asn Phe	
20 25	
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attactgtct tcagaaaact catgatgatc ctggccatga atg aaa agg ata aga Met Lys Arg Ile Arg	235
, met hys Arg Tie Arg -35	
aga aag aga aga aat gaa gtg acc atc cag cct ttc cca att aga ctt	283
Arg Lys Arg Arg Asn Glu Val Thr Ile Gln Pro Phe Pro Ile Arg Leu	
-30 -25 -20	
cct ctc ctt cca ccc ctc att tcc ttt ttg cac aca tta cag gtg gtg	331
Pro Leu Leu Pro Pro Leu Ile Ser Phe Leu His Thr Leu Gln Val Val	
-15 -10 -5 Egt tot gtg ata atg aaa ago ato aga aaa got ttt gta ott tgt ggt	270
Cys Ser Val Ile Met Lys Ser Ile Arg Lys Ala Phe Val Leu Cys Gly	379
1 5 10	
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aagaagagte tgttatgatg tgtaatacca atttctggag ggc atg gct gct ctc
                                                                       115
                                                 Met Ala Ala Leu
cga agt act cta aca tgg aca gaa gtc gtg ggc tgg tgg agt gtt gcg
                                                                       163
Arg Ser Thr Leu Thr Trp Thr Glu Val Val Gly Trp Trp Ser Val Ala
                                     -15
tcg ctg ctt agt gat gtg gca gca tgg tgg cca ccg cac tcc acc tca
                                                                      211
Ser Leu Leu Ser Asp Val Ala Ala Trp Trp Pro Pro His Ser Thr Ser
aca cgg gga ggg gta
                                                                      226
Thr Arg Gly Gly Val
    10
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                                                                      113
                                Met Asn Ala Leu Val Asp Gly Lys
cgg ctt asa krg tgc ata cgc tat ttc gat tct atc tca cta tat tct
                                                                      161
Arg Leu Xaa Xaa Cys Ile Arg Tyr Phe Asp Ser Ile Ser Leu Tyr Ser
                        -30
aag gca agt tta agt tgt tgt tta gtg tgt gtg ttt act tgt tca ttg
                                                                      209
Lys Ala Ser Leu Ser Cys Cys Leu Val Cys Val Phe Thr Cys Ser Leu
                    -15
                                        -10
cta gct ttc ttc agc cca tgc ac
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Teu Ala Phe Phe Ser Pro Cys
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<223> Von Heijne matrix
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<400> 302

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Met Thr Met Ala Val

-20

ggt gca gct gmy cam ctc ccc tgc tgc tgc cat ttr ctc acc tgc gtm 104 Gly Ala Ala Xaa Xaa Leu Pro Cys Cys Cys His Leu Leu Thr Cys Val

-10 -5

tcc agc ctt cgc amt gac att tac cca cat gg
Ser Ser Leu Arg Xaa Asp Ile Tyr Pro His

. <210> 303

<211> 175

<212> DNA

<213> Homo sapiens

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<221> CDS

<222> 73..174

<221> sig_peptide

<222> 73..147

<223> Von Heijne matrix
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seq SILLAALSRNISP/GQ

<400> 303

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-5

-10 cga aca tcc ccc gcg g Arg Thr Ser Pro Ala

175

5

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<212> DNA

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<221> sig peptide

<222> 402..470

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ttctttggag aaataccttt tccaaatcca atgggttgtc ttttttatt gttgatctta 180
agggttctta ggtgttctgg gtaccagttt cttgtgagat gtgtgacttg taaatacttt 240
cttccattct ccatgttgtc tttttattct cttgatggta ttctttgaaa tacaaaartk 300

tttatatttg acaaagttca gtttatttat ttatttattg ccattcgtgc ttttggtttt 360 gataatccat ttttwttgtt tttattttta tttacttaga g atg ggg tct ccc tat 416 Met Gly Ser Pro Tyr -20 gtt gcc cac gtt ggt ctt gaa ctc ttg acc tca agt gat cct ccc tcc 464 Val Ala His Val Gly Leu Glu Leu Leu Thr Ser Ser Asp Pro Pro Ser -15 -10 ttg gcc tcc caa gtg ctg gga ata cat tm 493 Leu Ala Ser Gln Val Leu Gly Ile His 1 <210> 305 <211> 214 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 79..213 <221> sig_peptide <222> 79..135 <223> Von Heijne matrix score 5.69999980926514 seq VCWLTLTLAHSHS/LT <400> 305 cacacacgca ccaaatacac acagasaccc tggccctcac tcacqcacav tctctcacac 60 tegtggacae acceecag atg cat ett tae act eat qta tge tgg ete act 111 Met His Leu Tyr Thr His Val Cys Trp Leu Thr -15 ctc aca ctg gca cac tca cac agc ttg acc cac acq cac aca ctc aca 159 Leu Thr Leu Ala His Ser His Ser Leu Thr His Thr His Thr Leu Thr -5 ccc agt cac aca cgt aca cac tca cat acg tgt gct tgc cta cac gca 207 Pro Ser His Thr Arg Thr His Ser His Thr Cys Ala Cys Leu His Ala 10 15 cac aag g 214 His Lys 25 <210> 306 <211> 458 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 306..458 <221> sig peptide <222> 306..350 <223> Von Heijne matrix score 5.69999980926514 seq LSLTFYHFPLCWG/HQ <221> misc feature <222> 286,448 <223> n=a, g, c or t

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cac cag gct gtg ccc acg tgg tgg saa rgc atc att caa cct tgt cac His Gln Ala Val Pro Thr Trp Trp Xaa Xaa Ile Ile Gln Pro Cys His 1 5 10 15	398
tgt gcc ctc tgc act tct gca gaa ggt gtg caa tca cat atc ata agt Cys Ala Leu Cys Thr Ser Ala Glu Gly Val Gln Ser His Ile Ile Ser 20 25 30	446
gna att tac aga Xaa Ile Tyr Arg 35	458
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gaactgetet gecaceette teetea atg aat gtg tta ate att gtt ttt gtt  Met Asn Val Leu Ile Ile Val Phe Val  -10  gea ttt get ttt ggg tte ytg gte atg aag tet ttg ett aag eea atg	
gaactgetet gecaccette teetea atg aat gtg tta ate att gtt ttt gtt  Met Asn Val Leu Ile Ile Val Phe Val  -10  gca ttt gct ttt ggg tte ytg gte atg aag tet ttg ett aag eea atg Ala Phe Ala Phe Gly Phe Leu Val Met Lys Ser Leu Leu Lys Pro Met	113
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gaactgetet gecaccette teetea atg aat gtg tta ate att gtt ttt gtt  Met Asn Val Leu Ile Ile Val Phe Val  -10  gca ttt gct ttt ggg tte ytg gte atg aag tet ttg ett aag eea atg Ala Phe Ala Phe Gly Phe Leu Val Met Lys Ser Leu Leu Lys Pro Met  -5  1  5  10  teg aga agg gtt ttt etg atg tta tet tet agg att ttt atg gtt tea Ser Arg Arg Val Phe Leu Met Leu Ser Ser Arg Ile Phe Met Val Ser	113
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gaactgetet gecaccette teetea atg aat gtg tta ate att gtt ttt gtt  Met Asn Val Leu Ile Ile Val Phe Val  -10  gca ttt gct ttt ggg tte ytg gte atg aag tet ttg ett aag eea atg Ala Phe Ala Phe Gly Phe Leu Val Met Lys Ser Leu Leu Lys Pro Met  -5  1  5  10  teg aga agg gtt ttt etg atg tta tet tet agg att ttt atg gtt tea Ser Arg Arg Val Phe Leu Met Leu Ser Ser Arg Ile Phe Met Val Ser	113
gaactgetet gecaccette teetea atg aat gtg tta ate att gtt ttt gtt  Met Asn Val Leu Ile Ile Val Phe Val  -10  gca ttt gct ttt ggg tte ytg gte atg aag tet ttg ett aag eea atg Ala Phe Ala Phe Gly Phe Leu Val Met Lys Ser Leu Leu Lys Pro Met  -5  1  5  10  teg aga agg gtt ttt etg atg tta tet tet agg att ttt atg gtt tea Ser Arg Arg Val Phe Leu Met Leu Ser Ser Arg Ile Phe Met Val Ser  15  20  25  ggt ett aga ttt aag tee ttg ate eat ett gag ttg att ttt gta tat Gly Leu Arg Phe Lys Ser Leu Ile His Leu Glu Leu Ile Phe Val Tyr  30  35  40	113 161 209 257
gaactgetet gecaccette teetea atg aat gtg tta ate att gtt ttt gtt  Met Asn Val Leu Ile Ile Val Phe Val  -10  gca ttt gct ttt ggg tte ytg gte atg aag tet ttg ett aag eea atg Ala Phe Ala Phe Gly Phe Leu Val Met Lys Ser Leu Leu Lys Pro Met  -5  1  5  10  teg aga agg gtt ttt etg atg tta tet tet agg att ttt atg gtt tea Ser Arg Arg Val Phe Leu Met Leu Ser Ser Arg Ile Phe Met Val Ser  15  20  25  ggt ett aga ttt aag tee ttg ate eat ett gag ttg att ttt gta tat Gly Leu Arg Phe Lys Ser Leu Ile His Leu Glu Leu Ile Phe Val Tyr  30  35  40  aag ttg aga gat gag gat eea gtt tea tte tte tae atg tgg ett gee	113 161 209
gaactgetet gecaccette teetea atg aat gtg tta ate att gtt ttt gtt  Met Asn Val Leu Ile Ile Val Phe Val  -10  gca ttt gct ttt ggg tte ytg gte atg aag tet ttg ett aag eea atg Ala Phe Ala Phe Gly Phe Leu Val Met Lys Ser Leu Leu Lys Pro Met  -5  1  5  10  teg aga agg gtt ttt etg atg tta tet tet agg att ttt atg gtt tea Ser Arg Arg Val Phe Leu Met Leu Ser Ser Arg Ile Phe Met Val Ser  15  20  25  ggt ett aga ttt aag tee ttg ate eat ett gag ttg att ttt gta tat Gly Leu Arg Phe Lys Ser Leu Ile His Leu Glu Leu Ile Phe Val Tyr  30  35  40	113 161 209 257
gaactgetet gecaccette teetea atg aat gtg tta ate att gtt ttt gtt  Met Asn Val Leu Ile Ile Val Phe Val  -10  gca ttt gct ttt ggg tte ytg gte atg aag tet ttg ett aag eea atg Ala Phe Ala Phe Gly Phe Leu Val Met Lys Ser Leu Leu Lys Pro Met  -5  1  5  10  tcg aga agg gtt ttt etg atg tta tet tet agg att ttt atg gtt tea Ser Arg Arg Val Phe Leu Met Leu Ser Ser Arg Ile Phe Met Val Ser  15  20  25  ggt ett aga ttt aag tee ttg ate eat ett gag ttg att ttt gta tat Gly Leu Arg Phe Lys Ser Leu Ile His Leu Glu Leu Ile Phe Val Tyr  30  35  40  aag ttg aga gat gag gat eea gtt tea tte tte tae atg tgg ett gee Lys Leu Arg Asp Glu Asp Pro Val Ser Phe Phe Tyr Met Trp Leu Ala  45  50  55  aat tat eec age ace att tgt tg	113 161 209 257
gaactgetet gecaccette teetea atg aat gtg tta ate att gtt ttt gtt  Met Asn Val Leu Ile Ile Val Phe Val  -10  gca ttt get ttt ggg tte ytg gte atg aag tet ttg ett aag eea atg Ala Phe Ala Phe Gly Phe Leu Val Met Lys Ser Leu Leu Lys Pro Met  -5  1  teg aga agg gtt ttt etg atg tta tet tet agg att ttt atg gtt tea  Ser Arg Arg Val Phe Leu Met Leu Ser Ser Arg Ile Phe Met Val Ser  15  20  25  ggt ett aga ttt aag tee ttg ate eat ett gag ttg att ttt gta tat Gly Leu Arg Phe Lys Ser Leu Ile His Leu Glu Leu Ile Phe Val Tyr  30  35  40  aag ttg aga gat gag gat eea gtt tea tte tte tee atg tgg ett gee  Lys Leu Arg Asp Glu Asp Pro Val Ser Phe Phe Tyr Met Trp Leu Ala  45  50  55  aat tat eee age ace att tgt tg  Asn Tyr Pro Ser Thr Ile Cys	113 161 209 257 305
gaactgetet gecaccette teetea atg aat gtg tta ate att gtt ttt gtt  Met Asn Val Leu Ile Ile Val Phe Val  -10  gca ttt gct ttt ggg tte ytg gte atg aag tet ttg ett aag eea atg Ala Phe Ala Phe Gly Phe Leu Val Met Lys Ser Leu Leu Lys Pro Met  -5  1  5  10  tcg aga agg gtt ttt etg atg tta tet tet agg att ttt atg gtt tea Ser Arg Arg Val Phe Leu Met Leu Ser Ser Arg Ile Phe Met Val Ser  15  20  25  ggt ett aga ttt aag tee ttg ate eat ett gag ttg att ttt gta tat Gly Leu Arg Phe Lys Ser Leu Ile His Leu Glu Leu Ile Phe Val Tyr  30  35  40  aag ttg aga gat gag gat eea gtt tea tte tte tae atg tgg ett gee Lys Leu Arg Asp Glu Asp Pro Val Ser Phe Phe Tyr Met Trp Leu Ala  45  50  55  aat tat eec age ace att tgt tg	113 161 209 257 305
gaactgctct gccacccttc tectca atg aat gtg tta atc att gtt ttt gtt  Met Asn Val Leu Ile Ile Val Phe Val  -10  gca ttt gct ttt ggg ttc ytg gtc atg aag tct ttg ctt aag cca atg Ala Phe Ala Phe Gly Phe Leu Val Met Lys Ser Leu Leu Lys Pro Met  -5	113 161 209 257 305
gaactgetet gecaccette teetea atg aat gtg tta ate att gtt ttt gtt  Met Asn Val Leu Ile Ile Val Phe Val  ———————————————————————————————————	113 161 209 257 305
gaactgetet gecaecette teetea atg aat gtg tta atc att gtt ttt gtt  Met Asn Val Leu Ile Ile Val Phe Val  -10  gca ttt gct ttt ggg tte ytg gte atg aag tet ttg ett aag cea atg Ala Phe Ala Phe Gly Phe Leu Val Met Lys Ser Leu Leu Lys Pro Met  -5	113 161 209 257 305
gaactgetet gecaccette teetea atg aat gtg tta atc att gtt ttt gtt  Met Asn Val Leu Ile Ile Val Phe Val  ———————————————————————————————————	113 161 209 257 305
gaactgctct gccacccttc tectca atg aat gtg tta atc att gtt ttt gtt  Met Asn Val Leu Ile Ile Val Phe Val  —10  gca ttt gct ttt ggg ttc ytg gtc atg aag tct ttg ctt aag cca atg Ala Phe Ala Phe Gly Phe Leu Val Met Lys Ser Leu Leu Lys Pro Met  —5 10  tcg aga agg gtt ttt ctg atg tta tct tct agg att ttt atg gtt tca Ser Arg Arg Val Phe Leu Met Leu Ser Ser Arg Ile Phe Met Val Ser  ———————————————————————————————————	113 161 209 257 305
gaactgetet gecaccette teetea atg aat gtg tta atc att gtt ttt gtt  Met Asn Val Leu Ile Ile Val Phe Val  ———————————————————————————————————	113 161 209 257 305
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173

Asn

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gaggwtcact tgagctkagg agttcaagga tgcagtsacc tgtgattgca ccactgcatt
                                                                      180
ccagcttgga caacagagtg agaccctgtc ttaaaattta aattttktgt yttwtggtag
                                                                      240
ag atg ggg tet ege eet gtt tee gak get ggt ete gaa ete etg gee
                                                                      287
   Met Gly Ser Arg Pro Val Ser Xaa Ala Gly Leu Glu Leu Leu Ala
                           ~15
teg age aat tet tet gee ttg eee tte eaa tgt tet ggg att aca gge
                                                                      335
Ser Ser Asn Ser Ser Ala Leu Pro Phe Gln Cys Ser Gly Ile Thr Gly
atg agc crc cac acc cta gcg g
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Met Ser Xaa His Thr Leu Ala
                15
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tgtgggggrg artatagtka cgaaaaagrk tattgtttcc cataatgcct ggtattgtat
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                                                                      240
taagtacttt gcatacagta gggcatttca ttgtcccagt gatcctcctg caaagtaggt
acaattatot toaatttaca aatgaggaaa ccaagototo ttoaagotga taagatgotg
                                                                      300
                                                                      360
aactgagatt tgaaccaagt coetetgeee ctaagageee etaceeetag etgetactat
abgotgtaco catotaagot tigigaaata roottgitoo acigoagaga ag aig big
                                                                      13.3
                                                           Met Leu
tgt cac cta tct cta gta ttt ctt ggc ktt ggg cag ttc tgg agt caa
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Cys His Leu Ser Leu Val Phe Leu Gly Xaa Gly Gln Phe Trp Ser Gln
    -10
aat g
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                                                                      120
tteggagega cettggeegt tggeetgace atetttgtge tgtetgtegt cactateate
                                                                      180
atotgottca cotgotoctg otgotgoott tacaagacgt googoogacc acgtooggtt
                                                                      240
gtcaccacca ccacatccac cactgtggtg c atg nnc ctt atc ctc agc ctc
                                                                      292
                                    Met Xaa Leu Ile Leu Ser Leu
                                    -15
caa gtg tgc cgc cca gct acc ctg gac caa gct acc agg gct acc aca
                                                                      340
Gln Val Cys Arg Pro Ala Thr Leu Asp Gln Ala Thr Arg Ala Thr Thr
            -5
cca tgc cgc cta cgg g
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Pro Cys Arg Leu Arg
    10
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                                            Met Cys His Arg Arg
                                                    -35
tgg etg cac cta hoa acchogt cat eta ggt tit aag ood ege atc cat
                                                                      3.02
Trp Leu His Leu Ser Thr Arg His Leu Gly Phe Lys Pro Arg Ile His
                             -25
tac gta ttt gtc tta atg ctg .tcc ctc ccc ttg ccc ccc acc ccc caa
                                                                      150
Tyr Val Phe Val Leu Met Leu Ser Leu Pro Leu Pro Pro Thr Pro Gln
    -15
                                             -5
cag gcc ctc ggg
                                                                      162
Gln Ala Leu Gly
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      seq FVIFPAALLLCWG/GL
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                                                                       60
atgaccagaa ategttteec tetetggggg gtteetgttt aatatgaaag teetettaae
                                                                      120
aagcgtggac agaggaagtt ttaggtttga tttgaacttc atgtacatga catatttcat
                                                                      180
tttttttttt tccctcacaa atttcaaccc aggccacttg tttgcagaga ctgccaaacc
                                                                      240
ttccattgct gcttccaaga tactcctgga atctgagatt accttttatc ctcttg atg
                                                                      299
gac cat gtt gtt att ttt gtc att ttc cct gca gct ctt ctg ctt tgc
                                                                      347
Asp His Val Val Ile Phe Val Ile Phe Pro Ala Ala Leu Leu Cys
            -15
                                -10
                                                     -5
tgg gga gga ete ate eee eta tge ate ate tae eee eeg ata get gae 🤄
                                                                      395
Trp Gly Gly Leu Ile Pro Leu Cys Ile Ile Tyr Pro Pro Ile Ala Asp
                                             10
aca gtt ggg
                                                                      404
Thr Val Gly
15
<210> 317
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<222> 359..448
<221> sig_peptide
<222> 359..433
<223> Von Heijne matrix
      score 5.59999990463257
      seq LIIILXFDIYSLA/FI
<221> misc feature
<222> 323,410
<223> n=a, g, c or t
<400> 317
tatgtetttt gaatttgtga tgtacatatt aacagtagat taagttgaaa taataaaats
                                                                       60
tgtattgttt afgafttatc agttatatga tgagtagaat atagtctatt gtggscmagt
                                                                      120
gtgtatatat aacataaaca atacattaac ccuattttgt ytgaaaatta ttttgggacc
                                                                      180
tagtagettt ettggteaca acettteaaa caaacaaatt ttttttaaat taattttte
                                                                      240
ccttaataaa gaaaacaatt cctcaatgtg taatagcaaa taccttttaa caggtcatat
                                                                      300
atcatcaatg ctttctttga aancgtactg atgcttacaa gatgctttac gagtaaag
                                                                      358
atg ctt aca aat ctt ttc ttt caa gta gct cat cct ctg atc att att
                                                                      406
Met Leu Thr Asn Leu Phe Phe Gln Val Ala His Pro Leu Ile Ile Ile
                    -20
                                         -15
                                                             -10
ctg ntg ttt gat atc tac tcc cta gca ttt atc cat gac gtg gg
                                                                      450
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177	
Leu Xaa Phe Asp Ile Tyr Ser Leu Ala Phe Ile His Asp Val -5 1 5	
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<pre>&lt;221&gt; sig_peptide &lt;222&gt; 313354 &lt;223&gt; Von Heijne matrix</pre>	
<pre>&lt;400&gt; 318 aatcgaaaac agcaaatcac acaactgtta aaaatatttt vtgtttacaa agccaagcca aaattttatg ttttcctccc caaactttga tataaacact aacatttttt agcatgtata aacatcatta ttaaccagtg tcctattaaa actccttttc tatgatagaa tgtctgttrc ttttaggtgg ataaggccta gatgattggc ctctaccagc atcctcatct ctgtccctga tgcccagctt carcctcgct cctgyatgct ggaccgcttc agtghagctc tcagacttgc tctgtgtctc ac atg cty ttt ggc tta cgt gga atg ctc cca ctc acc cag</pre>	60 120 180 240 300 351
caa gct ccc att cct cat tta aga tgt aaa ttg agt gtc acc tc Gln Ala Pro Ile Pro His Leu Arg Cys Lys Leu Ser Val Thr 1 5 10	395
<210> 319 <211> 257 <212> DNA <213> Homo sapiens	
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<pre>&lt;400&gt; 319 catctgtgtg tgcgtgtgt atg cgt gtg tgt atg cgt ctg tgt gca tgt gtg</pre>	52
tat gcg tgt gtg tgc gca tca gtg tct gca tgt gtg tat rtg tgt gta  Tyr Ala Cys Val Cys Ala Ser Val Ser Ala Cys Val Tyr Xaa Cys Val  -10 -5 1 5	100
tgt atg tst gtg cgc gcg cat ctg tgt gtg tgc atg tgt gta tgt atg Cys Met Xaa Val Arg Ala His Leu Cys Val Cys Met Cys Val Cys Met 10 15 20	148
tgt gtg cat ctc tgt gtg tgc atg tgt gta tgt gtg tgt gca tct gtg Cys Val His Leu Cys Val Cys Met Cys Val Cys Val Cys Ala Ser Val 25 30 35	196
tgt gtg tgc atg tgt gca tgc gtg tgt atg tgt gtg tgc gtg cgt gca Cys Val Cys Met Cys Ala Cys Val Cys Met Cys Val Cys Val Arg Ala 40 45 50	244
tct gtg tgt gtg c	257

178 Ser Val Cys Val <210> 320 <211> 325 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 256..324 <221> sig peptide <222> 256..318 <223> Von Heijne matrix score 5.59999990463257 seq LIANLVLFISIAA/LR <400> 320 accacgette etecaagtee cagegaacee gegtgeaace tgteectaaa aaagecaaag 60 cagteactet ttacetecca etttecetee teccageett tggcaaceae taatetaett 120 teegtgtata tggatttace tatteaggae attteatatg teetttggtg aetggettet 180 ttcactttgc acaatgtttt taaggttcat tcctgtcata gtgtgtgtca gtacgaaccc 240 ctccttaacc atcta atg gtt atc acc tct aat agt tat ctc ata gcc aat Met Val Ile Thr Ser Asn Ser Tyr Leu Ile Ala Asn -20 ctt gtt tta ttt ata tct atc gcc gcc ctc cgg g 325 Leu Val Leu Phe Ile Ser Ile Ala Ala Leu Arg <210> 321 <211> 201 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 31..201 <221> sig peptide <222> 31..183 <223> Von Heijne matrix score 5.5 seq LSLHASLVTKAFS/IN <400> 321 catcacaaga acccagagtg gaattctggg atg gaa gag ctg gac aga aag tgg 54 Met Glu Glu Leu Asp Arg Lys Trp -50 aga gag aag gtc ctc cca gcg gca aag cta att aaa agg aga aac ctg 102 Arg Glu Lys Val Leu Pro Ala Ala Lys Leu Ile Lys Arg Arg Asn Leu -40 -35 ttt tcc aca tgc act cct caa tat ggy aca cat gct gct ttc ttg tca 150 Phe Ser Thr Cys Thr Pro Gln Typ Gly Thr His Ala Ala Phe Leu Ser -25 -20 tta cat god toa ctt gid acc aaa goa tti toa atc aat too tgg gag 198 Leu His Ala Ser Leu Val Thr Lys Ala Phe Ser Ile Asn Ser Trp Glu

1

201

<210> 322 <211> 159

tgg

Trp

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 <222> 77..157
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 <223> Von Heijne matrix
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       seq PLLLCPLSSGSPC/PR
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                                                                     60
 tggaagtggg ctgggg atg gtg tcg ggg gcc caa gct ccc agc tcc caa agg
                                                                    112
                  Met Val Ser Gly Ala Gln Ala Pro Ser Ser Gln Arg
                  -25
                                      -20
 ccc ctg ctt cta tgc cct ttg agc tca ggt agc ccc tgc ccc cgg gg
                                                                    159
 Pro Leu Leu Cys Pro Leu Ser Ser Gly Ser Pro Cys Pro Arg
             -10
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 <222> 325..420
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 <222> 325..405
 <223> Von Heijne matrix
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      score 5.5
       seq SFLPSLLSSFLLS/LP
 <221> misc_feature
<222> 117
 <223> n=a, g, c or t
 <400> 323
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                                                                    120
 cgaaacaagc ctgagcaata gaagtagatg tggaaataac ttcgqtttct caaqqcaaat
                                                                    180
 actitigatag gaacaaacaa ccgtttagat atagaagatg tgatacattc ctttaaaaag
                                                                    240
 aatttgacct tatgtcattg taggcacacc tcatatttca attattcata tagtttttct
                                                                    300
 tgagcaattg ctggtttaag aata atg tca tgt ctt ttg cgt gct tat atc
                                                                    351
                           Met Ser Cys Leu Leu Arg Ala Tyr Ile
                                   -25
                                                       -20
 399
 Ile Trp Ile Phe Pro Ser Phe Leu Pro Ser Leu Leu Ser Ser Phe Leu
             -15
                                -10
 ctt tcc ctc ccc cct tcc ggg
                                                                    420
 Lou Ser Leu Pro Pro Ser Gly
        1
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<222> 71..181 <221> sig_peptide <222> 71..166 <223> Von Heijne matrix score 5.5 seq TLALLSSDSVATG/SV <400> 326 aatttcgcgg cctagtgggg cgtacgggcc tcttttgaaa gcctgagtta cgatgtattg 60 agegegtegt atg egg eca gea eta agg tee tte tgg eac tee tet ggt 109 Met Arg Pro Ala Leu Arg Ser Phe Trp His Ser Ser Gly -25 -30 gga eeg eee cea teg gee aca ett gee etg etc tee agt gat tet gta 157 Gly Pro Pro Pro Ser Ala Thr Leu Ala Leu Leu Ser Ser Asp Ser Val -15 -10 get act ggc tec gta gtc teg egg 181 Ala Thr Gly Ser Val Val Ser Arg 1 <210> 327 <211> 185 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 39..185 <221> sig_peptide <222> 39..116 <223> Von Heijne matrix score 5.5 seq LFSGWLVWWGSRS/SQ <221> misc_feature <222> 143,145,175 <223> n=a, g, c or t <400> 327 caaagacgca ctacttagta cagagaggtt ttgaatac atg ctc tgt gca tgc aag Met Leu Cys Ala Cys Lys -25 gca cgt ggg gtg atg ctg ctg ttc tca ggg tgg ttg gtt tgg tgg 104 Ala Arg Gly Val Met Leu Leu Phe Ser Gly Trp Leu Val Trp Trp -15 ggc agt agg tcc tca cag two ctc aga atg cct gag agn tna gta agt 152 Gly Ser Arg Ser Ser Gln Xaa Leu Arg Met Pro Glu Xaa Xaa Val Ser ggg gag ggt cga agc gat cdv dng cca cat ggg 185 Gly Glu Gly Arg Ser Asp Xaa Xaa Pro His Gly <210> 328 <211> 210 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 57..209

<221> sig_peptide <222> 57..182 <223> Von Heijne matrix score 5.5

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<400> 328

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ccc cgc g 210 Pro Arg

<210> 329

<211> 318

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 149..316

<221> sig_peptide

<222> 149..286

<223> Von Heijne matrix score 5.5 seq ILLISTLFYSLLS/GS

gatatttcag ttcctgattg taaatacctc ctaagcctga agcttctgtt actagccatt 120 gtgrgcttca gtktcttcak yckgcaaa atg ggc ata ata car kct att ctt 172 Met Gly Ile Ile Gln Xaa Ile Leu -45 gcc aca tca agg gat tgt tat tcc ttt aaa aaa aaa cca ata cca aag 220 Ala Thr Ser Arg Asp Cys Tyr Ser Phe Lys Lys Lys Pro Ile Pro Lys -35 -30 aag oot aca atg ttg goo tta goo aaa att otg ttg att toa acg ttg 268 Lys Pro Thr Met Leu Ala Leu Ala Lys Ile Leu Leu Ile Ser Thr Leu -15 ttt tat tca ctt cta tcg ggg agc cat gga aaa gra aat caa gac gtg 316 Phe Tyr Ser Leu Leu Ser Gly Ser His Gly Lys Xaa Asn Gln Asp Val

60

318

acacttaacc catctgtttt ctctaatgca cgacagattc ctttcagaca ggacaactgt

gg

<210> 330

<211> 223

<2125 DMA

<213> Homo sapiens

<220>

<221> CDS

<222> 135..221

<221> sig_peptide

<400> 332

<222> 135203 <223> Von Heijne matrix score 5.5 seq LPFVCLLLRNVYS/DL	
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-20 -15  ttt gta tgt ctt ctt ttg aga aat gtc tat tca gat ctt ttg ccc aat  Phe Val Cys Leu Leu Arg Asn Val Tyr Ser Asp Leu Leu Pro Asn -10 -5 1 5  cgg gg  Arg	218 223
<210> 331 <211> 362 <212> DNA <213> Homo sapiens	
<220> <221> CDS <222> 272361	
<221> sig_peptide <222> 272343 <223> Von Heijne matrix score 5.5 seq LIVVLVCISLVII/DD	
<400> 331 aatggacacc taggttgctt ccatatctga gctattgtga ataatgctgc aatgaacatg ggagtggaga catctcctaa gcatactgat ttcagttcct ttgggtatat acccagaagt gggatcatgt ggtaatcttg tttttacttt tttgaggaac ctccatacca ttatccatga tggctatagt aatttacatt cataccagca gtgcacaagg gtctcctttt ctgtatacac ttgccaacac ttgttatctt tcatttttt g atg cta gcc att cta aca ggt  Met Leu Ala Ile Leu Thr Gly -20	60 120 180 240 292
ggg agg tgg tat ctc ata gtg gtt tta gtt tgc att tcc ttg gtg att Gly Arg Trp Tyr Leu Ile Val Val Leu Val Cys Ile Ser Leu Val Ile -15 -10 -5	340
att gat gat gag cac ggg g Ile Asp Asp Asp Glu His Gly 1 5	362
<210> 332 <211> 89 <212> DNA <213> Homo sapiens	
<220> <221> CDS 22 3487	
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184	
cccagaccgg tcttgaactc ctggcctcaa ctg atg ctc ctg cct ctg ggt ctc Met Leu Leu Pro Leu Gly Leu	54
-10 aaa gtg ctg gga tta cag gcg aga ggc acc acg ct Lys Val Leu Gly Leu Gln Ala Arg Gly Thr Thr -5	89
<210> 333 <211> 399 <212> DNA <213> Homo sapiens	
<220> <221> CDS <222> 255398	
<pre>&lt;221&gt; sig_peptide &lt;222&gt; 255338 &lt;223&gt; Von Heijne matrix     score 5.5     seq PTLLVMWLSPQMA/SS</pre>	
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-25 -20 ggg gtc act ccc act ttg tta gtg atg tgg tta tct cct cag atg gcc Gly Val Thr Pro Thr Leu Leu Val Met Trp Leu Ser Pro Gln Met Ala -15 -10 -5	338
agt tcg ccc tct cag gct cct ggg atg gaa ccc tgc gcc tct ggg atc Ser Ser Pro Ser Gln Ala Pro Gly Met Glu Pro Cys Ala Ser Gly Ile 1 5 10 15 tca caa cgg gca a Ser Gln Arg Ala	386 399
20 <210> 334 <211> 188 <212> DNA <213> Homo sapiens	
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acc cct agc tca tat ccc atg ccc agt cat aaa cat gta tcc cta tgt Thr Pro Ser Ser Tyr Pro Met Pro Ser His Lys His Val Ser Leu Cys -25 -20 -15	101
ctt cta acg gtt gca gtt tta gtt ctt aca ttt aag tct tta att cat	149

188

100

280

300

185 Leu Leu Thr Val Ala Val Leu Val Leu Thr Phe Lys Ser Leu Ile His

-10 -5 1 ttt gag tda att ttt gca tat gag ata ggg gtc cag ggg

Phe Glu Xaa Ile Phe Ala Tyr Glu Ile Gly Val Gln Gly

15

<210> 335

<211> 115

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 23..115

<221> sig_peptide

<222> 23..94

<223> Von Heijne matrix score 5.5 seq CPSLLSPISPSQA/CP

<400> 335

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tcc tgt ccc tcc ctc ctc agc ccc atc tcc cca tcc cag gcc tgt cct Ser Cys Pro Ser Leu Leu Ser Pro Ile Ser Pro Ser Gln Ala Cys Pro -5 -10

gag ccc ctc ctt ggg 115 Glu Pro Leu Leu Gly

<210> 336

<211> 300

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 197..298

<221> sig_peptide

<222> 197..268

<223> Von Heijne matrix

score 5.5

seq IMFVCMCVCVCVC/VY

<400> 336

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-20

atg itt gta tgt atg tgc gtg tgt gtg tgt gtg tgt gtg tat cga etg Met Phe Val Cys Met Cys Val Cys Val Cys Val Cys Val Tyr Arg Leu -10

-5

ttt tct tcc tcc tcc ccg gg Phe Ser Ser Ser Pro

<210> 337

<211> 307

186

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   <220>
   <221> CDS
   <222> 5..307
  <221> sig peptide
  <222> 5..277
   <223> Von Heijne matrix
        score 5.5
        seq RVLLGAGIPPVSS/AP
  <400> 337
  caca atg aag teg aet gtt teg teg agg gaa gtg gee aee gtt gat aaa
                                                                          49
       Met Lys Ser Thr Val Ser Ser Arg Glu Val Ala Thr Val Asp Lys
                                -85
  atg aaa aga cgc cat gca gaa tac tgt gca cag ggt ctc cag aga ttt
                                                                          97
  Met Lys Arg Arg His Ala Glu Tyr Cys Ala Gln Gly Leu Gln Arg Phe
      -75
                           -70
                                               -65
  aaa gcc caa ctt tct caa gat acc ctt ccc cav cat cca cat ctg gag
                                                                         145
  Lys Ala Gln Leu Ser Gln Asp Thr Leu Pro Xaa His Pro His Leu Glu
                      -55
                                           -50
  awa gag aag ggg ctt gaa ggc ttg gag gaa aat gtg cct cta aag gga
                                                                         193
  Xaa Glu Lys Gly Leu Glu Gly Leu Glu Glu Asn Val Pro Leu Lys Gly
                   -40
                                       -35
  gag aaa cct gga gaa ggg ggt cca gag tct cct aag aag aga agg
                                                                         241
  Glu Lys Pro Gly Glu Gly Pro Glu Ser Pro Lys Lys Arg Arg Arg
              -25
                                   -20
  gtg ctt ctc gga gcg ggc atc cca cca gta agc tca gct ccc agg aga
                                                                        .289
  Val Leu Leu Gly Ala Gly Ile Pro Pro Val Ser Ser Ala Pro Arg Arg
          -10
                               -5
  cag agc cag cag gca aca
                                                                        307
  Gln Ser Gln Gln Ala Thr
                      10
  <210> 338
  <211> 123
  <212> DNA
  <213> Homo sapiens
  <220>
  <221> CDS
  <222> 16..123
  <221> sig_peptide
  <222> 16..75
  <223> Von Heijne matrix
        score 5.5
        seq VHLFFFFFXETGS/RS
  <400> 338
  ttaattaaac tgtgg atg cac aac agt tgt aga cct gtg cac ctt ttt ttc
                                                                         51
                   Met His Asn Ser Cys Arg Pro Val His Leu Phe Phe
                                       -15
ttt ttt ttt yct gag aca ggt tot egt tot aat yet tgy ctg gag tsc
                                                                         99
 Phe Phe Phe Xaa Glu Thr Gly Ser Arg Ser Asn Xaa Trp Leu Glu Xaa
 agt ggt gcg atc ata gct aac tcc
                                                                        123
 Ser Gly Ala Ile Ile Ala Asn Ser
     10
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<210> 339

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187
<211> 451
<212> DNA
<213> Homo sapiens
<220>
<221> CDS
<222> 318..449
<221> sig peptide
<222> 318..443
<223> Von Heijne matrix
      score 5.40000009536743
      seq TFRLYLSLPVSQA/GP
<221> misc feature
<222> 310..311,394
<223> n=a, g, c or t
<400> 339
gtcacaaaag gagcactaag agcctgcttt actttcttcc tcagttgagt cgtggggaca
                                                                        60
gcttgaagga gccaacctca attgcagaga gcagccgtca ccccagctac cgctcagage
                                                                       120
ccagcttgga accagagage ttecgttete ctacetttgg caaaagtttt caettegate
                                                                       180
cactatecag tggeteaege teeteeagee teaagteage ceagggeaea ggetttgage
                                                                       240
tgggccagtt gcaatccatt cgttcagagg gcaccacctc cacctcctaa taagagcctg
                                                                       300
gccaaccagn nacgcaa atg gaa gcc tat ctt aat gac agc ttg ctc aca
                                                                       350
                   Met Glu Ala Tyr Leu Asn Asp Ser Leu Leu Thr
cct tca gac agc cct gat ttt gag tca gtg cag gca ggg cct gna gcc
                                                                       398
Pro Ser Asp Ser Pro Asp Phe Glu Ser Val Gln Ala Gly Pro Xaa Ala
                        -25
                                             -20
aga ccc acc ttt agg cta tac ctc tcc ctt cct gtc agc cag gct ggc
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                                                                      114
                                     Met Pro Ala Leu Gly Pro Ala
ctt ctc cag ggc tct ctg kgc cgv gtg ggt cct cac cct cca gcs cct
                                                                      162
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188	
Leu Leu Gln Gly Ser Leu Xaa Arg Val Gly Pro His Pro Pro Ala Pro -5 5	
tcc acc aac tgc att cac tcc caa tgg cac gta tct gca gca csk ggc Ser Thr Asn Cys Ile His Ser Gln Trp His Val Ser Ala Ala Xaa Gly 10 15 20 25	210
aag gga ccc cac ctc agg cac cct ctr sct ggg nns tac caa ctt cct Lys Gly Pro His Leu Arg His Pro Leu Xaa Gly Xaa Tyr Gln Leu Pro 30 35 40	258
gtt cca gct gag ccc tgg gct gca gct gga ggc cac agt gtc cac c Val Pro Ala Glu Pro Trp Ala Ala Ala Gly Gly His Ser Val His 45 50 55	304
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gtc ccc tcg ctc tgt tgt tcc agc tat gt Val Pro Ser Leu Cys Cys Ser Ser Tyr -5 1	379
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Ile Leu Leu Ala Ser Gln Ala Gly Cys Leu Arg Ser Phe Leu Gly	_

PCT/IB99/00712

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raa	atg	ttt	gcc	tca	CCC	agg	aga	tgg	agc	tct	ncg	aaq	qcc	ttc	tct	22
	Met	Phe	Ala	Ser	Pro	Arg	Arg	Trp	Ser	Ser	Xaa	Lys	Ala	Phe	Ser	
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Gly	Gln	Arg	Thr	Leu	Leu	Ser	Ala	Ile	Leu	Ser	Met	Leu	Ser	Leu	Ser	
			-15					-10					-5			
ttc	tcc	aca	aca	tcc	ctq	ctc	agc	aac	tac	taa	+++	ata		aca	cad	324
Phe	Ser	Thr	Thr	Ser	Leu	Leu	Ser	Asn	Tvr	Trn	Phe	U = I	Glv	The	Cla	225
		1				5			- / -		10	Val	GIY	1111	GIII	
aaa	ata	CCC	aag	CCC	cta	-	aaa	222	aat	ata		~~~				276
Lvs	Val	Pro	Lys	Dro	Len	Cve	Glu	Two	994	Tou	yca Na	31-	aay	cgc	בננ	372
15	• • • •		LJ J	110	20	Cys	Giu	цуѕ	Gry		Ата	Ala	гÀг	Cys		
	a t-cr	~~~	ata	t-0.0						25					30	
yac Nan	Mot	Dra	gtg	000	Tou	gat	gga	gat	acc	aac	aca	tcc	acc	cag	gag	420
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cctc	accc	cc c	tgga	g at	g cc	c at	a ca	it to	c qt	a tt	c ct	c ta	t ac	c cc	c acc	232
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Cys :	Phe	Glv	Āla	Ser	Leu	Pro	Pro	T.eu	222	7~~	Cov	Lou	Ton	999	cag	320
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aaa :				~++								25				
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Gly	ser	ser	Pne	тте	ser	Trp	GIA	Thr	Gln	Ala	Ala	Ile	Val	Glu	Leu	
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Xaa	Pro	His														
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                                     -55
gcc acc cag ttc ctc tcc tgg gat gca tcc agt gtt tac agt ttc tta
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Ala Thr Gln Phe Leu Ser Trp Asp Ala Ser Ser Val Tyr Ser Phe Leu
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tat atc ctc tca gca aga gtt aat gta gac gta dgc agm tac att cgt
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Tyr Ile Leu Ser Ala Arg Val Asn Val Asp Val Xaa Xaa Tyr Ile Arg
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                             -25
gtg tac ata ctt gcc tgt gtg ttt ttc ctc tca cac ccc ctt ttt aad
                                                                       253
Val Tyr Ile Leu Ala Cys Val Phe Phe Leu Ser His Pro Leu Phe Xaa
                        -10
sra cca aat ggt agt gta tat tgt cnm cgt cat tct ccc cct tac ctt
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Xaa Pro Asn Gly Ser Val Tyr Cys Xaa Arg His Ser Pro Pro Tyr Leu
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                                                                       106
Leu Pro Leu Ser Pro Thr Lys Phe Leu Asn Val Phe Leu Gly Leu Phe
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                    -30
                                         -25
sto tat lat sit caa ttg gta igt stg ott att att tet lig git lig
                                                                      154
Leu Tyr Tyr Leu Gln Leu Val Cys Leu Leu Ile Ile Ser Leu Val Leu
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cct gat ctt gga cca agt tct gca cta aat cag aca ctc atg ttg ctg Pro Asp Leu Gly Pro Ser Ser Ala Leu Asn Gln Thr Leu Met Leu Leu -20 -15 -10	220
cgt gaa gtt tta gca tct cac gat tct tca gtk gta cca tta gat gct Arg Glu Val Leu Ala Ser His Asp Ser Ser Val Val Pro Leu Asp Ala -5 1 5 10	268
cgt caa gct gat ttt gtg cag ggg g Arg Gln Ala Asp Phe Val Gln Gly 15	293
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cag aac aca cac aac att kga gta cac cat ctt gtg tgg ctg tgg ttc Gln Asn Thr His Asn Ile Xaa Val His His Leu Val Trp Leu Trp Phe -20 -15 -10	222
gtg gtc ccc caa aca att aca atg ata aca cca aag atc act gaa cac Val Val Pro Gln Thr Ile Thr Met Ile Thr Pro Lys Ile Thr Glu His -5 5	270
age coe ste ete en out etr dir ete ete eve een tit gee eve tra	210

Arg Pro Xaa Ile Thr Asp Xaa Xaa Ile Met Xaa Thr Phe Glu Xaa Leu 10 15 20 gga gaa tta ccc a 331 Gly Glu Leu Pro <210> 355 <211> 93 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 2..91 <221> sig peptide <222> 2..55 <223> Von Heijne matrix score 5.30000019073486 seg ALYLCVCVCVCLI/AR <400> 355 t atg tgt ctv agt gta gct ttg tat tta tgt gtg tgt gtg tgt gta tqt 49 Met Cys Leu Ser Val Ala Leu Tyr Leu Cys Val Cys Val Cys Val Cys -15 -10 ctg att gca cgg gtg tac ttt tgt att tat gtg tgt gtg tgg tt 93 Leu Ile Ala Arg Val Tyr Phe Cys Ile Tyr Val Cys Val Trp 5 <210> 356 <211> 178 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 92..178 <221> sig peptide <222> 92..133 <223> Von Heijne matrix score 5.30000019073486 seq LHLLFGLFPVLWM/FL <400> 356 tgaccettgt ccagtetttt ccaggaaaaa catgeeetca agatgttttt ctatettgag 60 gaaatgatgg aaatgagata gttccaaggg t atg ctt cac ctt ctt ttt qqc 112 Met Leu His Leu Leu Phe Gly -10 tta ttt cct gtt ctt tgg atg ttt cta gtg tat ttc ttt ctt tct 160 Leu Phe Pro Val Leu Trp Met Phe Leu Val Tyr Phe Phe Leu Ser Ser -5 1 ttt ttt ttt ttt ttt 178 Phe Phe Phe Phe Phe <210> 357 <211> 107 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 40..105

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ttg gag cga ggg gat gag ccc tgg gtc ctg gat gtt cag ggc acc tct

198 Leu Glu Arg Gly Asp Glu Pro Trp Val Leu Asp Val Gln Gly Thr Ser 60 65 ggg aaa gag cac ctg aag aag tca aca gcc cag ctc ttg gga cca gaa 440 Gly Lys Glu His Leu Lys Lys Ser Thr Ala Gln Leu Leu Gly Pro Glu 80 ctg aag tac aag gag ttg ay 460 Leu Lys Tyr Lys Glu Leu 90 <210> 361 <211> 318 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 153..317 <221> sig peptide <222> 153..263 <223> Von Heijne matrix score 5.30000019073486 seq ALSSLCVSWGTSS/TV <400> 361 ctcttttccg gttaacgcgg cgtgagaagc catgagcagc aaagtctctc gcgacaccct 60 gtacgaggcg gtgcgggaag tcctgcacgg gaaccagcgc aasgccgcaa gttcctggag 120 acggtggagt tgcaggatca gcttgaagaa ct atg atc ccc aga agg aca agc 173 Met Ile Pro Arg Arg Thr Ser get tet egg gea eeg tea gte eee caa aac gea gge tta agt eea ete 221 Ala Ser Arg Ala Pro Ser Val Pro Gln Asn Ala Gly Leu Ser Pro Leu -30 -25 -20 ecc gee cta agt tet etg tgt gtg tee tgg ggg acc age age act gtg 269 Pro Ala Leu Ser Ser Leu Cys Val Ser Trp Gly Thr Ser Ser Thr Val -10 -5 acg agg cta agg ccg tgg ata tcc ccc aca tgg aca tcg agg gcg cgg g 318 Thr Arg Leu Arg Pro Trp Ile Ser Pro Thr Trp Thr Ser Arg Ala Arg 5 10 <210> 362 <211> 360 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 192..359 <221> sig_peptide <222> 192..233 <223> Von Heijne matrix score 5.30000019073486 seq VCIFCFLTSKAFP/NP <221> misc_feature <222> 277 <223> n=a, g, c or t <400> 362

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199

199	
aatactttca cagtttttca tagcagaaat ttataaatta atgaagccca ctttatactt ttatttcttt t atg gtt tgc atc ttt tgt ttc tta act tcg aaa gct ttt Met Val Cys Ile Phe Cys Phe Leu Thr Ser Lys Ala Phe -10 -5	180 230
cct aac cct aga tca cag gat ttt ctc tta gat ttc tct agg cat tnt Pro Asn Pro Arg Ser Gln Asp Phe Leu Leu Asp Phe Ser Arg His Xaa  1 5 10 15	278
ata ggt tta ggt ttc aca ttt agg tcc gca atg cat ttt gaa aac ttc Ile Gly Leu Gly Phe Thr Phe Arg Ser Ala Met His Phe Glu Asn Phe 20 25 30	326
cgt ctg waa ggt ttg ggt caa gat tcc ctt tgt c Arg Leu Xaa Gly Leu Gly Gln Asp Ser Leu Cys 35 40	360
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-5 1 5  <210> 364 <211> 242 <212> DNA <213> Homo sapiens	
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<222> 71,73

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aagtggtgag nmncaagcag ccagtggtag gaagggtcat caagtcagtt gtcagaaacc
                                                                      120
teactk atg tea etg twt ahg eta tgt gae eet gae eta gtt eet tge
      Met Ser Leu Xaa Xaa Leu Cys Asp Pro Asp Leu Val Pro Cys
                   -20
                                       -15
cct ctc ttg atc tca gtt gct tta tct gta aaa ttt cac att tkt cag
                                                                      216
Pro Leu Leu Ile Ser Val Ala Leu Ser Val Lys Phe His Ile Xaa Gln
              - 5
caa gtc aac ctt cca tgt tcc tct ca
                                                                      242
Gln Val Asn Leu Pro Cys Ser Ser
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                                                                       48
      Met Met Ile Leu Ile Leu Glu His Ile Val Thr Xaa
                       -35
                                           -30
aaa aga aac ccc aaa cct gtt aca gtc cct gct ttt ctg csc cct tgc
                                                                       96
Lys Arg Asn Pro Lys Pro Val Thr Val Pro Ala Phe Leu Xaa Pro Cys
                    -20
                                        -15
ttg act tet tte tet tgt ket gga gea tet tte tet etk ttw ggt gdg
                                                                      144
Leu Thr Ser Phe Ser Cys Xaa Gly Ala Ser Phe Ser Leu Xaa Gly Xaa
                -5
aga agg ggt tgg caa cat ggc agc tgc tgc tcc acc att ccc tta ttt
                                                                      192
Arg Arg Gly Trp Gln His Gly Ser Cys Cys Ser Thr Ile Pro Leu Phe
csa act cta aat tcc ctt ggg cag gga ctc att ggc cca gcc tac ata
                                                                      240
Xaa Thr Leu Asn Ser Leu Gly Gln Gly Leu Ile Gly Pro Ala Tyr Ile
ggt gcd gg
                                                                      248
Gly Ala
40
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                                                                      120
aaacaqtaqa atattattca gctataaaaa agaacagagt acacttagca aactaagaat
                                                                      180
agaaggaact tecteaatet gataaaggae ateeatgaaa aaceeaceae taatgteata
                                                                      240
cttaatcatq aaaaaccgaa tgcttttctc ctaagatagg aaaaagacaa gt atg tct
                                                                      298
                                                           Met Ser
act cat gcc atc tct att cta ctt tgt att ggt gct tct agc cag ggc
                                                                      346
Thr His Ala Ile Ser Ile Leu Leu Cys Ile Gly Ala Ser Ser Gln Gly
                -10
                                                                      351
agg gg
Arg
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       Met Glu Glu Glu Thr Glu Glu Val Gly Gly Arg Ser Ser
           -30
                               -25
                                                    -20
egg aaa aat gea gee ace gte aac gee gee tee etg eea eeg tge tte
                                                                       96
Arg Lys Asn Ala Ala Thr Val Asn Ala Ala Ser Leu Pro Pro Cys Phe
                            -10
ggg gta aaa agc tgc cgt tgc cgt cgg tgc agt tgc cgt cgc tgc ctc
                                                                      144
Gly Val Lys Ser Cys Arg Cys Arg Arg Cys Ser Cys Arg Arg Cys Leu
cta tac ttc tct tgg cct cgg gga agg att tcc cca ccg gtg gga caa
                                                                      192
Leu Tyr Phe Ser Trp Pro Arg Gly Arg Ile Ser Pro Pro Val Gly Gln
                                                                      208
tgt gcg ggg agg gga t
Cys Ala Gly Arg Gly
            35
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tcg g Ser 1													97
ggg g													145
gtt 1 Val 2													193
acg of Thr (													241
999 ( Gly 1 45													289
cat (													337
cac ( His )													385
ccc f				_			_		_				433
cac ( His ]			_	С									446
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                                                                        56
                                           Met Leu Leu Ala Val Ser
                                                        -10
ctg tcc ctt gtc tct aat tgt aac ttt gta ctc act gac caa ctt ttc
                                                                       104
Leu Ser Leu Val Ser Asn Cys Asn Phe Val Leu Thr Asp Gln Leu Phe
                             1
cct gcc cct gcs tcc ctc atc ccc gaa g
                                                                       132
Pro Ala Pro Ala Ser Leu Ile Pro Glu
<210> 371
<211> 127
<212> DNA
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                                                                        48
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    Met Asn Gln Asp Phe Asn Pro Glu Ile Glu Ala Ser Pro Gln Val
                     -25
                                         -20
aag act ggg gtt ttc ttg ttt tca att att ggg agt ttt gga ttt cca
                                                                        96
Lys Thr Gly Val Phe Leu Phe Ser Ile Ile Gly Ser Phe Gly Phe Pro
                 -10
                                     -5
gga atg tgc aat tgt aaa aac cca gcc cgg g
                                                                       127
Gly Met Cys Asn Cys Lys Asn Pro Ala Arg
<210> 372
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204 score 5.19999980926514

seq IVSSLFSWLLSLT/SV <221> misc_feature <222> 119 <223> n=a, g, c or t <400> 372 taaaaatctt ttatgttcta cccactcctt cctcgttccc tctccccact cctccctccc 60 cccatcttaa gcccatggca acccctgatc tttttactgt ctccatcgtt ttgccttbnc 120 caga atg cca tgt agt tgg agt cat ata gta agt agc ctt ttc agt tgg 169 Met Pro Cys Ser Trp Ser His Ile Val Ser Ser Leu Phe Ser Trp -15 ctt ctt tca ctt acc agt gtg ccc ggg 196 Leu Leu Ser Leu Thr Ser Val Pro Gly <210> 373 <211> 148 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 56..148 <221> sig_peptide <222> 56..139 <223> Von Heijne matrix score 5.19999980926514 seq PVLSCCCLTAGRA/RL <400> 373 actitettea cacceaggae geagggtgee getgeeggee acagaaacce caaga atg 58 ttt ttc ttt ggc tat tca gag gac atc tat tgt gtg tca ggc cct gtg 106 Phe Phe Gly Tyr Ser Glu Asp Ile Tyr Cys Val Ser Gly Pro Val -25 -20 ctg agc tgt tgt tgc ctg aca gca gga aga gcg cgg ctc tgg 148 Leu Ser Cys Cys Cys Leu Thr Ala Gly Arg Ala Arg Leu Trp -10 -5 <210> 374 <211> 200 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 26..199 <221> sig_peptide <222> 26..73 <223> Von Heijne matrix score 5.19999580926514 seq AALICPWSSQVPS/SP <400> 374 ctagggagga ctcaatgctc tttgt atg cct tat gca gcg ctg atc tgt ccc 52 Met Pro Tyr Ala Ala Leu Ile Cys Pro

-15 -10 tgg agt tcc cag gtt ccc agc tcc ccc cct gca agc ctt gaa gcc tcc

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205	
Trp Ser Ser Gln Val Pro Ser Ser Pro Pro Ala Ser Leu Glu Ala Ser -5 1 5	
agc aac gtc tat ctc cag gag agc agg gca gcc tat gca agt gtt ccg 1 Ser Asn Val Tyr Leu Gln Glu Ser Arg Ala Ala Tyr Ala Ser Val Pro 10 15 20 25	.48
gca gga cca gaa gtg gcc act caa cac acg tcc tca cca gtc acc cct 1 Ala Gly Pro Glu Val Ala Thr Gln His Thr Ser Ser Pro Val Thr Pro 30 35 40	.96
atg g Met	00
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	.05
	.12
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-20 -15 -10  tcc tcg gcg gct ccg agt cgt gcg agg cag ggg gcc c 16  Ser Ser Ala Ala Pro Ser Arg Ala Arg Gln Gly Ala -5 1	<b>4</b> 6
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attagtttaa ccattgtgga agatgatatg gcaattccac aaagacctaa agtcagraat
                                                                       120
tmcmattcaa cccagtaatc ccattactgg gtatatactc aaaggaatat aaattgttgt
                                                                       180
qttacaaaqa cacatgc atg cgt gtg ttc att gca gca ctg ttc aca ata
                                                                       230
                   Met Arg Val Phe Ile Ala Ala Leu Phe Thr Ile
                       -15
                                                                       261
gca gag aca tgg aat caa ccc aaa tgc cca g
Ala Glu Thr Trp Asn Gln Pro Lys Cys Pro
-5
                    1
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ag atg gca aaa ggc ctg agg gtg aat ctg ggc gag ctg gtt gag tcc
   Met Ala Lys Gly Leu Arg Val Asn Leu Gly Glu Leu Val Glu Ser
                                            -20
                        -25
                                                                       155
atg cqt ttq tqc ttc ctc tca qtc cac ttt cgc tta cga tgg ggc gac
Met Arg Leu Cys Phe Leu Ser Val His Phe Arg Leu Arg Trp Gly Asp
                     -10
tot tgt cca tcg tca cct cac cgg gaa act ttt cct gcc ggg cca gtt
                                                                       203
Ser Cys Pro Ser Ser Pro His Arg Glu Thr Phe Pro Ala Gly Pro Val
                                 10
                                                                       228
aat ggt ccc ctg tac cac ccc cgg g
Asn Gly Pro Leu Tyr His Pro Arg
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ttg awt tct tct kgr gtt tca aat gta aga ctg cta ctg tca cat ca

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                                                                       48
Met Pro His Pro Leu Ala Thr Ser Ala Phe Leu Arg Ser Ala Phe Pro
                        -20
                                             -15
ttt gtt tgt ctc acg ttt tgc gtg gga ggc ggt ccc ggg att tca ggg
                                                                       96
Phe Val Cys Leu Thr Phe Cys Val Gly Gly Pro Gly Ile Ser Gly
qtc tac cqq ctc ctt atq qcq aat qca acc cqa aqa qaq aqt qaq qta
                                                                      144
Val Tyr Arg Leu Leu Met Ala Asn Ala Thr Arg Arg Glu Ser Glu Val
ago etc ege ggg ttg ggc agg gac gga gag ggg gec ege geg act eca g
                                                                      193
Ser Leu Arg Gly Leu Gly Arg Asp Gly Glu Gly Ala Arg Ala Thr Pro
                            30
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                                                                       60
ctgctaattt cctcaggaca cattcccaga agtggaatta ccaagtcaaa gagcataaat
                                                                      120
actttagaga tacatgataa attgtgccag ctacctttcc aaaagagttg tactagttga
                                                                      180
ggtttctgcc agcagtat atg aca gtt ggg ctc cat att tta aga gat tca
                                                                      231
                    Met Thr Val Gly Leu His Ile Leu Arg Asp Ser
                                -20
cta atg gtg ttt ctc aac ctt ttt ttt aaac tgt gac cca cac agg
                                                                      279
Leu Met Val. Phe Leu Asn Leu Phe Phe Leu Asn Cys Asp Fro His Arg
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                                                                      281
99
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<213> Homo sapiens

PCT/IB99/00712 211

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His Gly Leu Arg Ser Ser Leu

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gtt gtc agt ggg ggt gtt ggt gac ttg acc aca aaa acc caa gag aat Val Val Ser Gly Val Gly Asp Leu Thr Thr Lys Thr Gln Glu Asn -30 -25 -20	105
ggg ctc tta cca gvc cty ctc tcc wkc ctk cac gga ctg ctc tat ggc Gly Leu Leu Pro Xaa Leu Leu Ser Xaa Leu His Gly Leu Leu Tyr Gly -15 -10 -5	153
agc cct gat gca gar ctc acg ggc ccg gat ccc tgg gat Ser Pro Asp Ala Glu Leu Thr Gly Pro Asp Pro Trp Asp 1 5 10	192
<210> 390 <211> 371 <212> DNA <213> Homo sapiens	
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tgc dtg cgc tcg ctc tct Cys Xaa Arg Ser Leu Ser	371

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                                                                    60
gcagtttatc tgcggtaact aaatgttagt tttgtaagta aaaggtactg ttattgacct
                                                                   120
cgaaagggct atagttcctt tgaacttaca gagaagagtt ccaaacaact atttctaacc
                                                                   180
aag atg gaa tat ggg tca gca aaa ttg tct tca ggt aga gtt ttc tac
                                                                   228
   Met Glu Tyr Gly Ser Ala Lys Leu Ser Ser Gly Arg Val Phe Tyr
                   -35
                                       -30
                                                                   276
ttg cca aga gac ttt ggc att gag agg aga gtt ctt gtt tgt ttt ttt
Leu Pro Arg Asp Phe Gly Ile Glu Arg Arg Val Leu Val Cys Phe Phe
               -20
                                   -15
aac tot gta toa ttt otg ttt ggt gto tot ara aaa aaa too gra caa
                                                                   324
Asn Ser Val Ser Phe Leu Phe Gly Val Ser Xaa Lys Lys Ser Xaa Gln
                                                                   328
tgg g
Trp
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<211> 303
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<221> sig_peptide
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ggctcactgt agccttgacc tcccaggctc aagcaatctt cctacctcag cctctcaggc
                                                                   180
agetgggaet acagacceae ageactaege etgaettatg attttatttt ttgtggagae
                                                                   240
                                                                   290
agggtcttac t atg ttg tct ggg ctt gtc tta aac tct tgg gcc tta gcc
            Met Leu Ser Gly Leu Val Leu Asn Ser Trp Ala Leu Ala
                        -10
                                                                   303
tac caa cta gct g
Tyr Gln Leu Ala
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seq FIAALFTVAKIWN/QP

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tgagcgctcg ccgtcttttg gcggcagcgg cgacgcgagg gctcccggcc gcccgcgtcc
                                                                   120
gctgggaatc tagcttctcc argamytgtg gtcgccccgt ccgctgtggc gggaaagcgg
                                                                   180
tccccagaac cgaccacacc gtggcaagag gacccagaac ccgaggacga aaacttgtat
                                                                   240
gagaaqaasc cagactccca tggknatgac aaggaccccg ttttggacgt ctggaac
                                                                   297
atg cga ctt gtc ttc ttc ktw ggc gks tcc atc atc ctg gtc ctt ggc
                                                                   345
Met Arg Leu Val Phe Phe Xaa Gly Xaa Ser Ile Ile Leu Val Leu Gly
                       -10
                                                                   366
age ace ttt gkg gcc tat ctg
Ser Thr Phe Xaa Ala Tyr Leu
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     seq SDFFLLFVSLSLS/PF
                                                                    53
agettggcat ataggeteaa atg tta tea tea gat ttt ttt ete ete ttt gte
                     Met Leu Ser Ser Asp Phe Phe Leu Leu Phe Val
                         -15
101
Ser Leu Ser Leu Ser Pro Phe Pro Phe Leu Phe Pro Pro Leu Phe
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tcc tgc ttt ctc tta ccc acc cgg g
Ser Cys Phe Leu Leu Pro Thr Arg
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Met Phe Ile Ala Ala Leu Phe -10												
aca gta gcc aag ata tgg aat caa cct aaa tgt cca tca acg gat gaa	222											
Thr Val Ala Lys Ile Trp Asn Gln Pro Lys Cys Pro Ser Thr Asp Glu -5 1 5												
tgg ata aat aaa atg tgg tac ata tac aca atg gag tac tat cca gac	270											
Trp Ile Asn Lys Met Trp Tyr Ile Tyr Thr Met Glu Tyr Tyr Pro Asp 10 20 25												
ata aaa aag aat gga att ctg aca ttt aag gca aca agg atg aac cgg	318											
Ile Lys Lys Asn Gly Ile Leu Thr Phe Lys Ala Thr Arg Met Asn Arg 30 35 40												
aag aca tta tg	329											
Lys Thr Leu												
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seq vedebevimeved/xv												
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4400- 306												
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Met Cys Val Cys Gly Cys Leu Cys Val Trp Met Cys Val Cys Gly -15 -10 -5												
wtt gtg tgt ata tac ata tgm gtg tat gtg tgt aca tgt gtg agg ggg	97											
Xaa Val Cys Ile Tyr Ile Xaa Val Tyr Val Cys Thr Cys Val Arg Gly												
1 5 10 15 ga	99											
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          Met Gln Phe Thr Val Leu Met Cys Pro Val Gln Trp Leu Leu
                  -20
                                       -15
gtg tat tca ccc agt tgt gca gcc acc atc aca gtc aat ttt aaa aca
                                                                        99
Val Tyr Ser Pro Ser Cys Ala Ala Thr Ile Thr Val Asn Phe Lys Thr
ttt tca tca ccc caa acc ggg
                                                                       120
Phe Ser Ser Pro Gln Thr Gly
<210> 400
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                                                                       120
gctkggggcc gccccagtag tgagacagtg gaagtaaacc ccatctgccg ttcccgtgcg
                                                                       180
                                                                       240
tagagaaaaa cgttgaccgc gaggctgggg aggagagttg cctctgagga agaagggcac
                                                                       300
agaganccaa aattagtttn gaaagcatcc tgatttggtg cccgaggcct ggaaagaaat
                                                                       356
ggcggctggg gtgcggcgga ggtaggggag gaaaacgttg g atg aga agg gcc tgg
                                               Met Arg Arg Ala Trp
                                                       -35
act cag gaa agg gaa ccg cgt ccg tgt gag ccc gct gag cgc gca gac
                                                                       404
Thr Gln Glu Arg Glu Pro Arg Pro Cys Glu Pro Ala Glu Arg Ala Asp
                                                                       452
cet gee cet gte tee tgt etg tet gea ggt etg ege gte tgt tgt tee
Pro Ala Pro Val Ser Cys Leu Ser Ala Gly Leu Arg Val Cys Cys Ser
    -15
cag cgc tct gc
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Gln Arg Ser
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Ser Gln Thr Leu Gln Arg Ser Ile Ser Phe Arg Ser Glu Ser Arg Pro

30

gac atc ctc gcc ccc cga ccc tgg tcc aga aat g Asp Ile Leu Ala Pro Arg Pro Trp Ser Arg Asn

15

330

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tgg aca aaa ttt tgt att tta att agt aca gca ttt cct tct tta ttg  Trp Thr Lys Phe Cys Ile Leu Ile Ser Thr Ala Phe Pro Ser Leu Leu  -15  -10  -5	4
aca cag att att ttc cct aaa tct att aca ttt gct ttc cag ttt ttc  Thr Gln Ile Ile Phe Pro Lys Ser Ile Thr Phe Ala Phe Gln Phe  1 5 10 15	2
tgg aac agg gaa aaa caa aaa aca aaa aca cca act ggg Trp Asn Arg Glu Lys Gln Lys Thr Lys Thr Pro Thr Gly 20 25	.1
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atg ctc gga ata ttt ttc aat gtc cat tcc gct gtg ttg att gag gac  Met Leu Gly Ile Phe Phe Asn Val His Ser Ala Val Leu Ile Glu Asp -10 -5 1 5	8
gtt ccc ttc acg gag aaa gat ttt gag aat ggc ccc cag aac ata tac Val Pro Phe Thr Glu Lys Asp Phe Glu Asn Gly Pro Cln Asn Ile Tyr 10 15 20	6
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Lys Ser Leu Lys Ile Cys Gly Leu Val Phe Gly Ile Leu Ala Leu Thr -10 -5 cta att gtc ctg ttt tgg ggg agc aag cac ttc tqq ccg gag gta ccc Leti Ille Val Leu Phe Trp Gly Ser Lys His Phe Trp Pro Glu Val Pro 15 aaa aaa gcc tat gac atg gag cac act acg gg 289 Lys Lys Ala Tyr Asp Met Glu His Thr Thr 20 25

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tte tee tgt ete gde cae etg agt age ega gae cae agg caa gea eta

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atg acc atg atc	gtt tcc ctt gct	gcg gtt gct tgg Ala Val Ala Trp -5		545
gtc cac aac ctg	ctt ctc acc tac	ctg ata gtg act Leu Ile Val Thr		593
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Phe Pro Pro 10

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227	
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aaaa atg ata cgt cct gtt tgt gaa ttg agc att ttt ttc acc tat gta	169
Met Ile Arg Pro Val Cys Glu Leu Ser Ile Phe Phe Thr Tyr Val -15 -10 -5	
cta gcc att tac ata tct cct tct gtg aat tgt ctg ttt ata tcc ttt Leu Ala Ile Tyr Ile Ser Pro Ser Val Asn Cys Leu Phe Ile Ser Phe  1 5 10	217
cct gcg gg Pro Ala 15	225
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ctc ctc ggg ctt cgt agc tct tgg ccc ggg gac cta cta agt gct cgg	104
Leu Leu Gly Leu Arg Ser Ser Trp Pro Gly Asp Leu Leu Ser Ala Arg -20 -15 -10	
ctc ttg tcc caa gag aag cgg gca gcg gaa acg cac ttt ggg ttt gag Leu Leu Ser Gin Glu Lys Arg Ala Ala Glu Thr His Phe Gly Phe Glu -5	152
act gtg tcg gaa gag gag aag agg ggg gac tta aca tca gtt gta agt Thr Val Ser Glu Glu Glu Lys Arg Gly Asp Leu Thr Ser Val Val Ser 10 15 20	200
cta gag tac cct gaa gtg caa tta cag ggt caa agg gtc tat gcm ttc Leu Glu Tyr Pro Glu Val Gln Leu Gln Gly Gln Arg Val Tyr Ala Phe	248

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                                                                       53
                                Met Glu Asn Leu Pro Phe Pro Leu
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                                                     -30
aaa tta ctt agt gcc tca tca cta aac acc ccc agc tcc aca cca tqq
                                                                      101
Lys Leu Ser Ala Ser Ser Leu Asn Thr Pro Ser Ser Thr Pro Trp
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gtg ttg gat atc ttc ctc acc ttg gtg ttt gcc ctg ggg ttc ttc ttc
                                                                      149
Val Leu Asp Ile Phe Leu Thr Leu Val Phe Ala Leu Gly Phe Phe Phe
    -10
                        -5
cta tta ctc ccc tac ttc tct tac ctc cgt tgt gac aac cca cca
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                               Met Cys Val Cys Val Phe Ala lie
                                            -10
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Phe Gly Val Arg Cys Cys Val Cys Val Arg Cys Ile
-5
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WO 99/53051 229 <211> 161 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 22..159 <221> sig_peptide <222> 22..153 <223> Von Heijne matrix score 4.80000019073486 seg XPCPLLFPGACFP/CP <400> 422 tcatttqqqt ttttatttaa t atg att tgc ata ttt tac tct aag att tcc 51 Met Ile Cys Ile Phe Tyr Ser Lys Ile Ser -40 atc tct gtc ggc tgt ggg agg aca gcc gag caa gtt gga tgt aaa 99 Ile Ser Val Gly Cys Gly Arg Thr Ala Ala Glu Gln Val Gly Cys Lys -25 cag agg tca ttt cac ckc ccy tgc cct ctg ctg ttt cct ggt gcd tgc 147 Gln Arg Ser Phe His Xaa Pro Cys Pro Leu Leu Phe Pro Gly Ala Cys -15 ttt ccc tgc cca ac 161 Phe Pro Cys Pro 1 <210> 423 <211> 420 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 333..419 <221> sig_peptide <222> 333..380 <223> Von Heijne matrix score 4.80000019073486 seq ICVSLMASDGASS/PV

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His Ser Ser Ser Xaa Xaa 10

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                                                                 240
ccaataaatg tgaac atg gga tct gtc acr gga gct gtc ctc aag acg cta
                                                                 291
                Met Gly Ser Val Thr Gly Ala Val Leu Lys Thr Leu
ctt ctg tta tct act caa aat tgg aac aga gtc gaa gct ggg aat tcc
                                                                 339
Leu Leu Ser Thr Gln Asn Trp Asn Arg Val Glu Ala Gly Asn Ser
                                      -25
                   -30
-35
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Tyr Asp Cys Asp Asp Pro Leu Val Ser Ala Leu Pro Gln Ala Ser Phe
               -15
                                  -10
age agt tet tee gag ete tee age agt cat agt eet gga ttt gea
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Ser Ser Ser Ser Glu Leu Ser Ser Ser His Ser Pro Gly Phe Ala
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Gln Asp Leu Val Val Lys Cys Ala Pro Pro Xaa Pro Phe Phe Leu Leu
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                   -20
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Phe Leu Phe Ser Ser Cys Asp Val Pro Val Pro Leu His Leu Leu Gln
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	)> 42 gtgag	-	taca	agtto	et ta	aaga	ıtggt	gtg	gtccg	ggag	tttg	gttco	ett o		atg Met	56
	_	_	_									gtg Val	-	ttg	ctg	104
	tca			_		caa			_		agt	gtt Val				152
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cagcgtctcg gtcacagagt tggagaaata gtgcagggac tcttcaggga gagcgttttc
ctcatcaaag caaactgcaa aatcgcttct gccggcgtgg acctg atg aga gtc ggt
Met Arg Val Gly
-30

	•
WO 99/53051	PCT/IB99/00712
ttt ggc cgc cta gga aac acc gag agt cac cta c Phe Gly Arg Leu Gly Asn Thr Glu Ser His Leu F 10 15	
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gtc tca gat gtg gtc agc ggg att ccc ttc aaa c Val Ser Asp Val Val Ser Gly Ile Pro Phe Lys I	

-10

132

51

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<400> 429 caaactgttg aaaagttaac tett atg tta ttt ata ttt tea gae ata gat

Met Leu Phe Ile Phe Ser Asp Ile Asp
-30 -25

tgg aag atg gac tta tgc ttt ttc tct ttc tct cct ccc tcc 99

Trp Lys Met Asp Leu Cys Phe Phe Ser Phe Ser Pro Phe Leu Pro Ser
-20 -15 -10

ctt cct ttg ttg gag gct gaa aga atg agg gtc agt gat caa ctt cag 147

Leu Pro Leu Leu Glu Ala Glu Arg Met Arg Val Ser Asp Gln Leu Gln
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tat acc act gga kac ggg

Tyr Thr Thr Gly Xaa Gly

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354

Pro Cys Gln Xaa Gly Leu 20

cca tgc cag tng gga ctg gg

-30

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accettgate geacteagaa accagaetta aaacetettt geagettete aggaeteage
                                                                      120
tggaaccaac gggcacagtt ggcaacacca tc atg aca tca caa cct gtt ccc
                                                                      173
                                    Met Thr Ser Gln Pro Val Pro
aat gag acc atc ata gtg ctc cca tca aat gtc atc aac ttc tcc caa
                                                                      221
Asn Glu Thr Ile Ile Val Leu Pro Ser Asn Val Ile Asn Phe Ser Gln
gca gag aaa ccc gaa ccc acc aac cag ggg cag gat agc ctg aag aaa
                                                                      269
Ala Glu Lys Pro Glu Pro Thr Asn Gln Gly Gln Asp Ser Leu Lys Lys
    -45
                                             -35
cat cta cac gca gaa atc aaa gtt att ggg act atc cag atc ttg tgt
                                                                    317
His Leu His Ala Glu Ile Lys Val Ile Gly Thr Ile Gln Ile Leu Cys
                    -25
                                         -20
ggc atg atg gta ttg agc ttg ggg atc att ttg gca tct gct tcc ttc
                                                                      365
Gly Met Met Val Leu Ser Leu Gly Ile Ile Leu Ala Ser Ala Ser Phe
                -10
tot coa aat tit acc caa gig act tot aca cig iig aac tot got tac
                                                                      413
Ser Pro Asn Phe Thr Gln Val Thr Ser Thr Leu Leu Asn Ser Ala Tyr
                            10
                                                15
cca ttc ata gga ccc ggg
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Pro Phe Ile Gly Pro Gly
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                                        Met Val Asp Glu Cys Leu
                                                            -35
                                        -40
aca gag cct gtg tgg gga agc aaa agg caa ggg tgt agt tca cag gca
                                                                      102
Thr Glu Pro Val Trp Gly Ser Lys Arg Gln Gly Cys Ser Ser Gln Ala
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235	
gaa gcg agc tgt gac att gtc agt gca gcg tgt aag tgt ggc tcc tca Glu Ala Ser Cys Asp Ile Val Ser Ala Ala Cys Lys Cys Gly Ser Ser -15 -10 -5	150
cag gcg gcc att gat tgt gag acc tca tct tgc tct gaa gat ttc ccg Gln Ala Ala Ile Asp Cys Glu Thr Ser Ser Cys Ser Glu Asp Phe Pro  1 5 10	198
gtg Val 15	201
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-10 -5 1	
gga tac ggc tct ctg atg gct cct tct agc cct acc cct tct ggg Gly Tyr Gly Ser Leu Met Ala Pro Ser Ser Pro Thr Pro Ser Gly 5 10 15	334
gga tac ggc tct ctg atg gct cct tct agc cct acc cct tct ggg Gly Tyr Gly Ser Leu Met Ala Pro Ser Ser Pro Thr Pro Ser Gly	334
gga tac ggc tct ctg atg gct cct tct agc cct acc cct tct ggg Gly Tyr Gly Ser Leu Met Ala Pro Ser Ser Pro Thr Pro Ser Gly 5 10 15  <210> 435 <211> 386 <212> DNA	334
gga tac ggc tct ctg atg gct cct tct agc cct acc cct tct ggg Gly Tyr Gly Ser Leu Met Ala Pro Ser Ser Pro Thr Pro Ser Gly 5 10 15  <210> 435 <211> 386 <212> DNA <213> Homo sapiens  <220> <221> CDS	334
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236

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Gln Val Cys Pro Arg Leu Cys Leu Gln Arg Xaa Val Gly Glu Leu Gln

239

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10
  mtg cnn nky cct gat gtg gga aca gct ctt ctc cca gat gtt aat aga
                                                                        322
  Xaa Xaa Xaa Pro Asp Val Gly Thr Ala Leu Leu Pro Asp Val Asn Arg
                                          25
  aca ago tgo aca aco tgg
                                                                        340
  Thr Ser Cys Thr Thr Trp
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  <211> 409
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                                                                        120
  gattggtgag ctgagtggag aagtgccata gagcggtgtt ttgccagagt gtctgcggat
                                                                        180
  tgctcatacc tgggaaggat tctttgtatg gttcccttag gctgagggag ggtatcagct
                                                                        240
  ttacagacct tgtgggatta caaaagggcc accacact cttcaaccaa t atg tgt
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                                                            Met Cys
  cta tct tgc att caa ggc tca ttc ttt gtt gaa att ttg cag ttg gtc
                                                                        345
  Leu Ser Cys Ile Gln Gly Ser Phe Phe Val Glu Ile Leu Gln Leu Val
                          -20
                                               -15
  act agg cta ttg tta tct cca tct caa agt aca cag aca cac aca cac
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  Thr Arg Leu Leu Ser Pro Ser Gln Ser Thr Gln Thr His Thr His
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  Thr His Thr His Thr
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tctqttcttt tataggaaga aaaaacatag ttatttttct tttatgatac aaaggtatgc 120

240	
tttctatgca agctggatac cagaccaaga ataataaatc acaatttcat aaggtttcta agacttgata ttatatgggg at atg acc att ttg agg gaa atg tnn nca tca Met Thr Ile Leu Arg Glu Met Xaa Xaa Ser -30 -25	180 232
ctt tat gta ctt gaa gct aag gat act gct atc tta ttg ctt gtt tna Leu Tyr Val Leu Glu Ala Lys Asp Thr Ala Ile Leu Leu Leu Val Xaa -20 -15 -10	280
gtg agc gat aag aat gaa cag cag ctt ggg agg ggc gtg g Val Ser Asp Lys Asn Glu Gln Gln Leu Gly Arg Gly Val -5	320
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tcc tgt ggg ttg cct gtt aag act ttg cca ttt atc tgt tgc aat ctt Ser Cys Gly Leu Pro Val Lys Thr Leu Pro Phe Ile Cys Cys Asn Leu -20 -15 -10	164
tat ttc ttg ctg ttt tgt agg agt tct ttt tta tat ttt gga tat gat Tyr Phe Leu Leu Phe Cys Arg Ser Ser Phe Leu Tyr Phe Gly Tyr Asp -5 1 5	212
ccc att aat act tac atg tat tac aat gtt ttc tcc cac tcg gg Pro Ile Asn Thr Tyr Met Tyr Tyr Asn Val Phe Ser His Ser 10 15 20	256
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agg ggt etg tgc ccc gcc cat ccc ggg gcc cct cct ttg ccc cgc ccc Arg Gly Leu Cys Pro Ala His Pro Gly Ala Pro Pro Leu Pro Arg Pro -40 -35 -30 -25	148
ccg gac cgc ctt ccc cat tca ttc tct cct acg ggg tgt ctc ctg hgc Pro Asp Arg Leu Pro His Ser Phe Ser Pro Thr Gly Cys Leu Leu Xaa -20 -15 -10	196
ccc ctt ctg gtc tcg tgt ttg ggg tct ctg ctt ccg gtc acc caa acc Pro Leu Leu Val Ser Cys Leu Gly Ser Leu Leu Pro Val Thr Gln Thr -5	244
ctg ggg tcc ttc agt gct ggt ccc tgc ttc agg acc ctc a Leu Gly Ser Phe Ser Ala Gly Pro Cys Phe Arg Thr Leu 10 15 20	284
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tgt cca ctt agc caa ttc cta cct att ctt tma agc ctc agt tcc agt Cys Pro Leu Ser Gln Phe Leu Pro Ile Leu Xaa Ser Leu Ser Ser	162
tgt cca ctt agc caa ttc cta cct att ctt tma agc ctc agt tcc agt	162 210
tgt cca ctt agc caa ttc cta cct att ctt tma agc ctc agt tcc agt Cys Pro Leu Ser Gln Phe Leu Pro Ile Leu Xaa Ser Leu Ser Ser Ser -20 -15 -10 gtc ccc tcg agg gca ggc agt gct ttc cca tct gcc cta ggt cca ctc Val Pro Ser Arg Ala Gly Ser Ala Phe Pro Ser Ala Leu Gly Pro Leu	
tgt cca ctt agc caa ttc cta cct att ctt tma agc ctc agt tcc agt  Cys Pro Leu Ser Gln Phe Leu Pro Ile Leu Xaa Ser Leu Ser Ser Ser  -20 -15 -10  gtc ccc tcg agg gca ggc agt gct ttc cca tct gcc cta ggt cca ctc  Val Pro Ser Arg Ala Gly Ser Ala Phe Pro Ser Ala Leu Gly Pro Leu  -5 1 5 10  tac cag cct cta ctt ggg ccc cca gca tgg  Tyr Gln Pro Leu Leu Gly Pro Pro Ala Trp	210
tgt cca ctt agc caa ttc cta cct att ctt tma agc ctc agt tcc agt  Cys Pro Leu Ser Gln Phe Leu Pro Ile Leu Xaa Ser Leu Ser Ser Ser  -20	210
tgt cca ctt agc caa ttc cta cct att ctt tma agc ctc agt tcc agt Cys Pro Leu Ser Gln Phe Leu Pro Ile Leu Xaa Ser Leu Ser Ser Ser -20 -15 -10  gtc ccc tcg agg gca ggc agt gct ttc cca tct gcc cta ggt cca ctc Val Pro Ser Arg Ala Gly Ser Ala Phe Pro Ser Ala Leu Gly Pro Leu -5 1 5 10  tac cag cct cta ctt ggg ccc cca gca tgg Tyr Gln Pro Leu Leu Gly Pro Pro Ala Trp 15 20  <210> 446 <211> 184 <212> DNA <213> Homo sapiens  <220> <221> CDS	210
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gct att tta ttt ccc aat tca gga tca tgc ttt gca ttt agt tgt cat Ala Ile Leu Phe Pro Asn Ser Gly Ser Cys Phe Ala Phe Ser Cys His -10 -5 1
gtc tcc ttt ttt ttt tt tt tt tt tt tt t t t t t
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ttc ttg tta agc cgg caa aac tgt gta agc aca gga kga cct tca tcc  146  Phe Leu Ser Arg Gln Asn Cys Val Ser Thr Gly Xaa Pro Ser Ser  15 20 25
aaa tct gat atc aac tca agg tct gga tct tgt tca ctg gca agg gg Lys Ser Asp Ile Asn Ser Arg Ser Gly Ser Cys Ser Leu Ala Arg 30 35 40
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15

270

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cag gtg ttg ttt tgt aat cga ag

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Gln Val Leu Phe Cys Asn Arg
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               Met Ser Tyr Phe Arg Cys Ile Phe Leu Ala Val Leu
                            -15
tca aaa atc agt tgg gct gta aat atg tgc agt ctt att tct ggg tcc
                                                                        97
Ser Lys Ile Ser Trp Ala Val Asn Met Cys Ser Leu Ile Ser Gly Ser
                                                              10
tcg gg
                                                                       102
Ser
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                                                                        55
                                       Met Leu Cys Ile Met Phe Gly
att gaa act aat gaa att acc aag atg aca atg tot ttt ott ttg ttt
                                                                       103
Ile Glu Thr Asn Glu Ile Thr Lys Met Thr Met Ser Phe Leu Leu Phe
        -25
                             -20
cta agt atc agt ttg ata act tta tat tat tcc tca gaa gca tgt ggg
                                                                       151
Leu Ser Ile Ser Leu Ile Thr Leu Tyr Tyr Ser Ser Glu Ala Cys Gly
    -10
                         -5
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Gln Val Pro Thr Tyr Gly Pro Tyr Gly Arg Cys Ala Pro Met Lys Ser 55 60 65 atc tcc agc agc ctc aag gag 311 Ile Ser Ser Ser Leu Lys Glu <210> 460 <211> 425 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 161..424 <221> sig_peptide <222> 161..418 <223> Von Heijne matrix score 4.59999990463257 seq AAAALCILILLXA/MY <400> 460 aggeogget gatgegeagg caatttatea tettgatete ceaetgagte agggagetet 60 cctqtcacca gtattgattt cagaggatgg actaaatttc ctaggatttc cattaagaat 120 175 taagaaaaaa gctctaagca cgcagggtag ccagacagac atg gat atg aga tgg Met Asp Met Arg Trp cac tgt gaa aac tcg cag acc aca gat gac atc ctt gtg gcc tca gca 223 His Cys Glu Asn Ser Gln Thr Thr Asp Asp Ile Leu Val Ala Ser Ala -70 -80 -75 gag tgt ccc agc gat gat gag gac att gac ccc tgt gag ccg agc tca 271 Glu Cys Pro Ser Asp Asp Glu Asp Ile Asp Pro Cys Glu Pro Ser Ser -55 -60 ggt ggg tta gcc aac cca acc cga gca ggc ggc aga gag ccg tat cca 319 Gly Gly Leu Ala Asn Pro Thr Arg Ala Gly Gly Arg Glu Pro Tyr Pro -35 ~45 -40 367 gge tea gea gaa gtg ate egg gag tee age age ace aeg ggt atg gte Gly Ser Ala Glu Val Ile Arg Glu Ser Ser Ser Thr Thr Gly Met Val -30 -25 gtt ggg ata gta gcc gct gcc gcc ctg tgc atc ctt atc ctc wat 415 Val Gly Ile Val Ala Ala Ala Leu Cys Ile Leu Ile Leu Leu Xaa -15 -10 425 gcc atg tac a Ala Met Tyr 1 <210> 461 <211>. 420 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 45..419 <221> sig_peptide <222> 45..104 <223> Von Heijne matrix score 4.59999990463257 seq PTLLTLCIGSVVS/SD

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tct gac ctg act cag gac cct gct gtg tct gtg gcc ttg gga cag aga Ser Asp Leu Thr Gln Asp Pro Ala Val Ser Val Ala Leu Gly Gln Arg  1 5 10 15	152
gtc agg atc aca tgc cag gga gac aac ctc gaa gag tat ttt gca agc Val Arg Ile Thr Cys Gln Gly Asp Asn Leu Glu Glu Tyr Phe Ala Ser 20 25 30	200
tgg tac cga cag agg ccc gga cag gcc cct gtc ctt gtc atc tat ggt Trp Tyr Arg Gln Arg Pro Gly Gln Ala Pro Val Leu Val Ile Tyr Gly 35 40 45	248
aaa aac aac cgg ccc tca ggg att cca gsc cgr ktc tct ggc tcc aag Lys Asn Asn Arg Pro Ser Gly Ile Pro Xaa Arg Xaa Ser Gly Ser Lys 50 55 60	296
tca ggc aat aca gct tta ttg acc atc gyc ggg gct cag gcg gag gat  Ser Gly Asn Thr Ala Leu Leu Thr Ile Xaa Gly Ala Gln Ala Glu Asp  65 70 75 80	344
gab gct gac tat tac tgt agt kat cgc gac cat act gat aat cgg tgg Xaa Ala Asp Tyr Tyr Cys Ser Xaa Arg Asp His Thr Asp Asn Arg Trp 85 90 95	392
gtg ttc ggc ggg ggg acc agg ctg aca g Val Phe Gly Gly Thr Arg Leu Thr 100 105	420
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ttt tac atg gkg att ctt acc tgc ttg atc ttc agg aac tca gaa ggg Phe Tyr Met Xaa Ile Leu Thr Cys Leu Ile Phe Arg Asn Ser Glu Gly -15 -10 -5	105
ttt cag att gyc cat gtc cag aaa caa cag tgt ctt ttc aaa aat gag Phe Gln Ile Xaa His Val Gln Lys Gln Gln Cys Leu Phe Lys Asn Glu 1 5 10 15	153
aaa gtg gtc gtg ggc tca tgc aac agg acc atc cag aac cag cag tgg Lys Val Val Gly Ser Cys Asn Arg Thr Ile Gln Asn Gln Gln Trp 20 25 30	201
atg tgg act gag gat gaa aag ete ett eat gtt aaa tet gea etg tge Met Trp Thr Glu Asp Glu Lys Leu Leu His Val Lys Ser Ala Leu Cys 35 40 45	249
ttg gcc at Leu Ala 50	257

251

seq LFRVLFSXTCALX/QD

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                                                                         60
 atg tgc gtg tgc gcg tgt gct ttg tgt gtg tgg ttg tgt gtt aaa tca
                                                                        108
 Met Cys Val Cys Ala Cys Ala Leu Cys Val Trp Leu Cys Val Lys Ser
                                                                        117
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 Cys Ser Ile
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                                            Met Ile Ser Asp Val Gln
                                                -20
 cac ctt ttc ata tac ttg tta gcc ttt tgt atg cct tcc ttg gag aaa
 His Leu Phe Ile Tyr Leu Leu Ala Phe Cys Met Pro Ser Leu Glu Lys
                     -10
                                                                        142
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 Cys Leu Tyr Gly Ser Leu Ala His Phe Phe Phe
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tecagtgeet gagagaaaac ggeetaateg aaaacgteeg eggeatacat ecattettaa
                                                                      120
aacttgagtg gctgcttttc tgggtggaaa agagcggtat cagacagggt gagcagtcgg
                                                                      180
ggaacggatg aacaaagact tgcaccgtgg ccctg atg cct ttg ttc cga gtt
                                                                      233
                                       Met Pro Leu Phe Arg Val
cta ttc agt tgw act tgt gcg ttg twa cag gac ttt aga atg cag ccc
                                                                      281
Leu Phe Ser Xaa Thr Cys Ala Leu Xaa Gln Asp Phe Arg Met Gln Pro
tgc ccc cca acc ccc aag g
                                                                      300
Cys Pro Pro Thr Pro Lys
        10
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<212> DNA
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aaaggataga aaagttacgt tgatggtgtg cccctcgata tctagaagat agcatagtcc
                                                                      120
atgcattctc agaaagatcc tatcc atg tgg tat gta gag atg tgg gtt tct
                                                                      172
                            Met Trp Tyr Val Glu Met Trp Val Ser
                                             -20
ttt ttt cta ctt ttt tat gtg ctt ctt ttt aga aac tta tac aca cac
                                                                      220
Phe Phe Leu Leu Phe Tyr Val Leu Leu Phe Arg Asn Leu Tyr Thr His
                    -10
aca cac cac act ggg
                                                                      235
Thr His His Thr Gly
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Met Ala Ala Arg Val Gly Ala Phe Leu Lys Asn Ala Trp Asp Lys Glu	
-30 -25 -20 -15	150
cca gtg ctg gtc gtg tcc ttc gtc gtc ggg ggc ctc ggc tgt aat dct	153
Pro Val Leu Val Val Ser Phe Val Val Gly Gly Leu Gly Cys Asn Xaa	
-10 -5 1	
gcc ccc att gag ccc cta ctt caa gta ctc cgt cat gat caa caa ggc	201
Ala Pro Ile Glu Pro Leu Leu Gln Val Leu Arg His Asp Gln Gln Gly	
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           -25
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Leu Ser Leu Ser Leu Ser Ala Ser Leu Ile Ile Ser Pro Ser Pro Ser
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gag atg tgc tgg ytg cgg gya tgg ggc cag atc ctc ctg cca gtt ttc 168 Met Cys Trp Leu Arg Xaa Trp Gly Gln Ile Leu Leu Pro Val Phe. -25 -20 cbn tcc ctc ttt ctc atc caa ttg ctt atc agc ttc tca gag aat ggt 216 Xaa Ser Leu Phe Leu Ile Gln Leu Leu Ile Ser Phe Ser Glu Asn Gly -5 1

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Leu Met Leu Xaa Xaa Tyr Trp Ser Cys Trp Ile Lys Ser Pro Pro Xaa -5 1 5

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151

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	ca gag aag aat gtg att	
=	hr Glu Lys Asn Val Ile	
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•	90 -85	-
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Phe Gly Ile Gly Tyr Val Thr Leu Leu Gln Ile His Ser Ile Tyr Ser	
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Ile Leu Thr Leu Asn Thr Val Phe Val Leu Ala Val Lys Leu Lys Trp	
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<222> 372..419

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                                                                       60
attetgacgg taactgtgta teagttggaa ttactgcact aactttgagg gecatactca
                                                                      120
aggactccaa taataaccaa gtcaatggcc ttagtggaaa tacaacaatt ccgtttagca
                                                                      180
gctgttgggc caactacaca gaccttactc cccttagaac aggaaaaaat tataagattg
                                                                      240
aatttatact ggataatgtt gttggggtag aatccagaac tttcagcctg ctggcagagt
                                                                      300
ctgtctctag cagtggcagc agcagcagca gcmacagcaa agcatcaact gtgggtacat
                                                                      360
atgcccagat a atg act gtm gta att agc tgt ctg gtt gga gaa tgt ggc
                                                                      410
             Met Thr Val Val Ile Ser Cys Leu Val Gly Glu Cys Gly
                             -10
tct tgg aaa t
                                                                      420
Ser Trp Lys
    1
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<400> 486
caaccatac atg tgc aca ctc aca gac aca cac act cac gtc caa gtg cac
                                                                        51
          Met Cys Thr Leu Thr Asp Thr His Thr His Val Gln Val His
                  -45
                                       -40
aag toa aaa oot tgo cag oto oto too oot oot coa coa rsc cat ggt
                                                                        99
Lys Ser Lys Pro Cys Gln Leu Leu Ser Pro Pro Pro Pro Xaa His Gly
                                 -25
ect ett ett etc ect atc ttt gge ett ett gtg ecc tet eag att tte
                                                                       147
Pro Leu Leu Pro Ile Phe Gly Leu Leu Val Pro Ser Gln Ile Phe
        -15
                            -10
age tet ett ete aat tet eta eat etg gge etg eet tee tte eea aag
                                                                       195
Ser Ser Leu Leu Asn Ser Leu His Leu Gly Leu Pro Ser Phe Pro Lys
                                         10
                                                             15
atg cca ctc atg att ttc ctc ccc cgc tgg g
                                                                       226
Met Pro Leu Met Ile Phe Leu Pro Arg Trp
                20
<210> 487
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<222> 221..454
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<222> 221..409 <223> Von Heijne matrix score 4.5 seq QILXSTLAMKIHS/QQ

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-55

gtg aag gat cta ttc tct ctg atg ata act tgg aca gtc cag atg aaa 331 Val Lys Asp Leu Phe Ser Leu Met Ile Thr Trp Thr Val Gln Met Lys -35 -40 ~30

ttg aca tca atg tgg atg aac ttg ata ccc ccg atg aag cag att ctt 379 Leu Thr Ser Met Trp Met Asn Leu Ile Pro Pro Met Lys Gln Ile Leu ~20 -15

tdg agt aca ctg gcc atg aag atc cac agc caa caa aga ttc tgg cca 427 Xaa Ser Thr Leu Ala Met Lys Ile His Ser Gln Gln Arg Phe Trp Pro -10 -5

aga gtc aga gtc tat tcc aga ata tac 454

Arq Val Arg Val Tyr Ser Arg Ile Tyr 10 15

<210> 488

<211> 329

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 253..327

<221> sig peptide

<222> 253..309

<223> Von Heijne matrix score 4.5

seq VLFLLNLFQKIEE/EE

## <400> 488

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-15

329

ttc caa aaa att gag gag gag gaa ctc ttc cct aat ga Phe Gln Lys Ile Glu Glu Glu Leu Phe Pro Asn -5 1

<210> 489

<211> 414

<212> DNA

<213> Homo sapiens

<220>

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<222> 149..412

<221> sig_peptide <222> 149..292

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265
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<222> 247..348
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<222> 247..333
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      seq ILLILQLLKXSLK/KC
<221> misc feature
<222> 323..324
<223> n=a, g, c or t
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ctataatttg tatctaaaat taggttttcc cttttaagtt gttaattttc tatggkttgt
                                                                      120
gctgcatgct ttcactttta ttagtactta cagccaaaga gatgggcaaa tgtctagaaa
                                                                      180
aattaatgtt ttgattcagg aatttgtgcc tagtgatggc ctccaataga gaattttcca
                                                                      240
gagaga atg aag act cag ttt cta agt tgg ggc aaa ttt agt ttt tgt
                                                                      288
       Met Lys Thr Gln Phe Leu Ser Trp Gly Lys Phe Ser Phe Cys
                       -25
ttt ggt att ctt ctt ata tta cag cta tta aaa bnn tct ctt aaa aaa
Phe Gly Ile Leu Leu Ile Leu Gln Leu Lys Xaa Ser Leu Lys Lys
-15
                                         -5
                                                                      348
tgc cgg cac ggg
Cys Arg His Gly
<210> 492
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<222> 5..124
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      score 4.5
      seq LRFILPSSWDCRC/AP
<400> 492
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                                                                       49
     Met Leu Pro Ala Val Ala Val Ser Glu Pro Val Val Leu Arg Phe
att ctg ccg agt tcc tgg gat tgc agg tgc gcg cca ctc ctg act
                                                                       97
Ile Leu Pro Ser Ser Trp Asp Cys Arg Cys Ala Prc Pro Leu Leu Thr
.-10 .
                     -5
ggu ttt lýc att ttt tgg ktg gag acg gg
                                                                      126
Gly Phe Cys Ile Phe Trp Xaa Glu Thr
            10
                                 15
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<210> 493
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266 <213> Homo sapiens <220> <221> CDS <222> 119..298 <221> sig peptide <222> 119..217 <223> Von Heijne matrix score 4.40000009536743 seq WLLMVAPRLPAGA/RD <400> 493 acactgcctg cctggtgcag cccatgtgac gggtcgagct ccgggccctg ctgtccctgg 60 cogggctate coagtggctt caggcacett etecagaeet acceagaaag atgecegg 118 atg gat cct gca gct ccg tgg ctt ttc tgg gaa gca gcg gcc cct gct 166 Met Asp Pro Ala Ala Pro Trp Leu Phe Trp Glu Ala Ala Ala Pro Ala -30 ctc aag aga ccc tgg ctc ctg atg gtg gcc cca agg ttg cca gct ggt 214 Leu Lys Arg Pro Trp Leu Leu Met Val Ala Pro Arg Leu Pro Ala Gly -10 get agg gac tea gga cag ttt cee aga aaa gge caa geg gge age eee 262 Ala Arg Asp Ser Gly Gln Phe Pro Arg Lys Gly Gln Ala Gly Ser Pro tee agg gge egg gtg agg aag etg ggg ggt geg gtg gg 300 Ser Arg Gly Arg Val Arg Lys Leu Gly Gly Ala Val 20 25 <210> 494 <211> 295 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 182..295 <221> sig peptide <222> 182..274 <223> Von Heijne matrix score 4.40000009536743 seq SRLXALLSPYAFT/LX <400> 494 tttatacaca cacacacaca cacactcata ttcattacat gtgtgtactt tctggttgct 60 tcagtaggac tttttctaggc ttctttggac tatgtgtgat attttacttc agggactgaa 120 tttcacaact gcctactatg caactttgtg attttcttga aagcacaakt actatatata 180 a atg aaa atg tcc acc ccc tcc ccg ctt tct aaa aaa gtg ctc aga aac 229 Met Lys Met Ser Thr Pro Ser Pro Leu Ser Lys Lys Val Leu Arg Asn -30 -25 cag gtc tca aga ttg rtt gcg ttg ctt tcc cca tac gct ttc act ctg 277 Gln Val Ser Arg Leu Xaa Ala Leu Leu Ser Pro Tyr Ala Phe Thr Leu -5 -10 1 sct cgt ctt gcc tca ggg 295 Xaa Arg Leu Ala Ser Gly

<210> 495

<211> 244

<212> DNA

<213> Homo sapiens

WO 99/53051 267 <221> CDS <222> 70..243 <221> sig_peptide <222> 70..114 <223> Von Heijne matrix score 4.40000009536743 seq RFLLLYATQQGQA/KA ggaagtegeg ttgtgeaggt tegtgeeegg etggegegge gtggttteae tgttacatge 60 cttgaagtg atg agg agg ttt ctg tta cta tat gct aca cag cag gga cag Met Arg Arg Phe Leu Leu Tyr Ala Thr Gln Gln Gly Gln -10 gca aag gcc atc gca gaa gaa atg tgt rag caa gct gtg gta cat gga 159 Ala Lys Ala Ile Ala Glu Glu Met Cys Xaa Gln Ala Val Val His Gly 10 ttt tct gca gat ctt cac tgt att agt gaa tcc gat aag gtc tcg gtg 207 Phe Ser Ala Asp Leu His Cys Ile Ser Glu Ser Asp Lys Val Ser Val 20 25 att cag aat aca cct act ttt gca acg ggg ggg cgg g 244 Ile Gln Asn Thr Pro Thr Phe Ala Thr Gly Gly Arg <210> 496 <211> 215 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 91..213 <221> sig_peptide <222> 91..171 <223> Von Heijne matrix score 4.40000009536743 seq FVNLNLCFAYTFA/LY atttaagtcc agagagcaag gtgattgcag tttctttgtt cggtttgctt attttttact 60 gcttatttct gtgtgcataa attcagcgac atg cta ata gac ata tgg tca atg 114 Met Leu Ile Asp Ile Trp Ser Met -25 gtg ctt aga gaa aat ctg ttt gta aac ctg aat ctc tgt ttt gcc tac 162 Val Leu Arg Glu Asn Leu Phe Val Asn Leu Asn Leu Cys Phe Ala Tyr -15 -10 aca ttt gca ttg tat tcc tgc cct gct cca act cgt tgt cct aga cca 210 Thr Phe Ala Leu Tyr Ser Cys Pro Ala Pro Thr Arg Cys Pro Arg Pro 1 10 tcc ag 215 Ser <210> 497

<211> 255

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<222> 36..254

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ctg cct gcc ttc ctc tgg ccg ctg ggg ata ccc tgg cct gat gga gag 149 Leu Pro Ala Phe Leu Trp Pro Leu Gly Ile Pro Trp Pro Asp Gly Glu 10 15

ggt cta aga cct tcc cgt ctt ctc cgg aca cgg gaa aac att acc cct 197 Gly Leu Arg Pro Ser Arg Leu Leu Arg Thr Arg Glu Asn Ile Thr Pro 30 25

ctc tct tta ttc gct atg ctg agt ggc agg ggt gcc ccg ctc ctg 245 Leu Ser Leu Phe Ala Met Leu Ser Gly Arg Glu Gly Ala Pro Leu Leu 45

255 gtc ccc ctg g Val Pro Leu

<210> 498

<211> 82

<212> DNA

<213> Homo sapiens

55

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<222> 23..82

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<400> 498

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tee ttg eeg geg gag ace eet aag caa ggg 82 Ser Leu Pro Ala Glu Thr Pro Lys Gln Gly

<210> 499

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<212> DNA

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269

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cyc gcc tgg ctg cag ccc agg tat agg aag aat gcg tat ctt ttc atc Xaa Ala Trp Leu Gln Pro Arg Tyr Arg Lys Asn Ala Tyr Leu Phe Ile -100 -95 -90	104
tat tac tta atc cag ttc tgt ggc cas tct tgg ata ttt gca aat atg Tyr Tyr Leu Ile Gln Phe Cys Gly Xaa Ser Trp Ile Phe Ala Asn Met -85 -70 -70	152
Thr Val Arg Phe Phe Ser Phe Gly Lys Asp Ser Met Val Asp Thr Phe -65 -60 -55	200
tat gct att gga ctt gtg atg cga ctt tgc caa tcc gta tct ctc ctg  Tyr Ala Ile Gly Leu Val Met Arg Leu Cys Gln Ser Val Ser Leu Leu  -50  -45  -40	248
gaa ctg ctg cac ata tat gtt ggc att gag tca aac cat ctt ctc cca Glu Leu Leu His Ile Tyr Val Gly Ile Glu Ser Asn His Leu Leu Pro -35 -30 -25	296
agg ttt ttg cag ctc aca gaa aga ata atc atc ctt ttt gtg gtg atc Arg Phe Leu Gln Leu Thr Glu Arg Ile Ile Leu Phe Val Val Ile -20 -15 -10	344
acc agt cga aga gga agt cca acg aga aat atg tgg tgt gtg tgt tat  Thr Ser Arg Arg Gly Ser Pro Thr Arg Asn Met Trp Cys Val Cys Tyr  -5 1 5 10	392
tcg tct ttg gat cta tgg ata tgg tta rgt aca ctt ata gca tgk tda  Ser Ser Leu Asp Leu Trp Ile Trp Leu Xaa Thr Leu Ile Ala Xaa Xaa  15 20 25	440
tca gtc ata gga ata tcc tat gct gtc ttg aca t Ser Val Ile Gly Ile Ser Tyr Ala Val Leu Thr 30 35	474
<210> 500 <211> 241 <212> DNA <213> Homo sapiens	÷
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	227
	24 î
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270

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gcagaagcct cgctgaatcc cagccagctg gttctaacct tccagaatcg caatcccttc

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180

271 totgagactg agcotgocat coactogoac gootttottt cagggotttt cggctgttgg 300 ctacactgat gtgacccccc tccctttttg ga atg atg ggg atc ttt ttg gtg 353 Met Met Gly Ile Phe Leu Val tat gtn gga ttt gtt ttc ttt tcc gtt tta tat gta caa caa ggg ctt 401 Tyr Val Gly Phe Val Phe Phe Ser Val Leu Tyr Val Gln Gln Gly Leu -15 tct tct caa gca 413 Ser Ser Gln Ala 1 <210> 503 <211> 167 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 26..166 <221> sig_peptide <222> 26..91 <223> Von Heijne matrix score 4.40000009536743 seq WVLDPALLLTCLT/FP <400> 503 gaateggaca acttaaagte tegat atg age ete gga ttg eat teg aae tee 52 Met Ser Leu Gly Leu His Ser Asn Ser -20 tgg gtt cta gac cca gct ctg cta cta act tgt ctg acc ttc ccc att 100 Trp Val Leu Asp Pro Ala Leu Leu Leu Thr Cys Leu Thr Phe Pro Ile -10 -5 tat aaa ctg ttg tgg gtg aga ggt ggg acw agg wga act ctr wgr gcv 148 Tyr Lys Leu Leu Trp Val Arg Gly Gly Thr Arg Xaa Thr Leu Xaa Ala 5 ctg cac tcg gcg cgg acg g 167 Leu His Ser Ala Arg Thr <210> 504 <211> 420 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 217..420 <221> sig_peptide <222> 217..396 <223> Von Heijne matrix score 4.40000009536743 seq MWVXCXFCFVLFC/FE <221> misc_feature <222> 47..48,355..369,373 <223> n=a, g, c or t <400> 504

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272	
gattgtaagg agaggcggtc ccggtgtcct cgggtcccag gtgattgtga agtgctgacc aattgccact ggacatactt gaaacaaaat aggaaa atg gca gca aac tct tca Met Ala Ala Asn Ser Ser -60 -55	180 234
gga caa ggt ttt caa aac aaa aat aga gtt gca atc ttg gca gaa ctg Gly Gln Gly Phe Gln Asn Lys Asn Arg Val Ala Ile Leu Ala Glu Leu -50 -45 -40	282
aca aag aga aaa gaa aac tac tta tgc aga acc agt ctt caa caa atc Thr Lys Arg Lys Glu Asn Tyr Leu Cys Arg Thr Ser Leu Gln Gln Ile -35 -30 -25	330
atc ctg gar cta ggt att gac act ata atg tgg gtt tnn tgt ntg ttt Ile Leu Glu Leu Gly Ile Asp Thr Ile Met Trp Val Xaa Cys Xaa Phe -20 -15 -10	378
tgt ttt gtt ttg ttt tgt ttt gag acg gag tct cgc cct gtc Cys Phe Val Leu Phe Cys Phe Glu Thr Glu Ser Arg Pro Val -5 1 5	420
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274

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277 -30 cta ctg gga acg gaa cag aaa caa aaa agg atg gga aat ctg aag 284 Leu Leu Gly Thr Glu Gln Lys Gln Lys Lys Arg Met Gly Asn Leu Lys -25 -20 ctg cta ttt ctt att ctg atc tta ata gca gga tac agg g 324 Leu Leu Phe Leu Ile Leu Ile Leu Ile Ala Gly Tyr Arg -5 <210> 514 <211> 303 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 212..301 <221> sig_peptide <222> 212..292 <223> Von Heijne matrix score 4.40000009536743 seq SALMLPLGCAVRT/RM <400> 514 tttccctcac tctctgcctc ccccatcgca ccccacagga gggtttccct cactctctgc 60 ctccccatc gcaacccaca ggagggtttc cctcactctg cctcctccaw cgcaccccca 120 kggaggtgtt ttccctcact ggttctgttg gtggcggtgg cagcaatccg agtcacatgg 180 caccagagta tgtcacgggt ggcggatctg a atg ggg ctg cag agc ctc aca 232 Met Gly Leu Gln Ser Leu Thr -25 ctt cca gtg tct tgc agc cct tct gcc ctg atg ctt ccc ttg gga tgt 280 Leu Pro Val Ser Cys Ser Pro Ser Ala Leu Met Leu Pro Leu Gly Cys -10 -15 303 gct gtc cgc acg cgc atg ctt ga Ala Val Arg Thr Arg Met Leu 1 <210> 515 <211> 455 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 342..455 <221> sig_peptide <222> 342..434 <223> Von Heijne matrix score 4.40000009536743 seq LTTLESLAGSVXS/EQ <400> 515 tcatatctgg waatggcaaa cagggatgaa aatcgattat gttttggaga ctccttttgg 60 acatgtatca gtgtgttgat ttgcacaaac caataaaagc cctacattti ttggaaatgg 120 atcoctagat ttcaagcatg tataatcact caaagtggat atgatcacag gcattcttct 180 ctigagetea geaaaaetat geetaeeaae aeegaagaga agteaaagat ttttatyaaa 240 aaaaattgca gatgatgttg gtgagataat aggatatgag caatgaaccc ttgggtgggg 300 ttccagggca cttaaattgc ctcgtgtctt gagtccttaa g atg gac tca aac aaa 356 Met Asp Ser Asn Lys -30 aaa tta gta tta tca ata aca ggt aat act gtg tgg att cta aca aca 404

Lys Leu Val Leu Ser Ile Thr Gly Asn Thr Val Trp Ile Leu Thr Thr

278 -25 -20 -15 tta gaa tca tta gct ggc agt gtc aam tct gaa caa gat ttg tca gct 452 Leu Glu Ser Leu Ala Gly Ser Val Xaa Ser Glu Gln Asp Leu Ser Ala -10 tat 455 Tyr <210> 516 <211> 360 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 196..360 <221> sig_peptide <222> 196..336 <223> Von Heijne matrix score 4.40000009536743 seg SFXXCLFLXLXXS/EM <221> misc_feature <222> 330..332 <223> n=a, g, c or t <400> 516 aagagcgttg ggcagatata gtctgtagat atttttgaaa cgtctttggg tttgtcccat 60 ttggggtttg ctcagcttct tgaatctgta ggttttgggg atcccccamc ctgcaaattt 120 ggtgatattt ttgctcttat ttctkcaagt gaacttgaaa tcccaccctg ttgqttttct 180 cettetaaga etetg atg acg tgt atg tta gee tgt agg tgt agt ete amg 231 Met Thr Cys Met Leu Ala Cys Arg Cys Ser Leu Xaa -45 ggt ccc caa gat ttt cgt ttc tgc tct gtc ttt tct ctg ttg ctc aag 279 Gly Pro Gln Asp Phe Arg Phe Cys Ser Val Phe Ser Leu Leu Lys -30 -25 ttg ggt aat ttc tat ttt tct ttt wct dtc tgt ctw ttt ctw dta ctd 327 Leu Gly Asn Phe Tyr Phe Ser Phe Xaa Xaa Cys Leu Phe Leu Xaa Leu -10 wyn nnt tct gag atg gag tcm cac tct ttc agc 360 Xaa Xaa Ser Glu Met Glu Ser His Ser Phe Ser <210> 517 <211> 453 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 113..451 <221> sig_peptide <222> 113..307 <223> Von Heijne matrix score 4.40000009536743 seq FIEAALLIHGSAC/VY <400> 517 attttcctgg gcgggaacag caaaatggcg ccagaactag tggcgggctg aggacgccgt 60 accectegga aggeageest geggteestt tgeegeeegt teeeteeegg as atg gag 118

Met Glu

279	
-65	
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acc aag aac tgg gag gtg gac gtg gcg gcc cag ctg ggc gag tat ctg Thr Lys Asn Trp Glu Val Asp Val Ala Ala Gln Leu Gly Glu Tyr Leu -45 -40 -35	214
gag gag ctg gat cag atc tgc att tct ttt gac gaa ggc aag acc aca Glu Glu Leu Asp Gln Ile Cys Ile Ser Phe Asp Glu Gly Lys Thr Thr -30 -25 -20	262
atg aac ttc att gag gca gcg ttg ttg atc cat ggc tct gcc tgc gtc Met Asn Phe Ile Glu Ala Ala Leu Leu Ile His Gly Ser Ala Cys Val -15 -10 -5 1	310
tac agt aag aag gtg gaa tac ctc tac tca ctc gtc tac cag gcc ctt Tyr Ser Lys Lys Val Glu Tyr Leu Tyr Ser Leu Val Tyr Gln Ala Leu 5 10 15	358
gat ttc atc tct gga aag agg cgg gcc aag cag ctc tct tcg gtg cag Asp Phe Ile Ser Gly Lys Arg Arg Ala Lys Gln Leu Ser Ser Val Gln 20 25 30	406
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Met Phe Val Ser Tyr Leu Ile Leu Thr Leu Leu His Val Gln Thr -15 -10 -5	
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280

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282

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<212> DNA

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tcc tc	t act	ctc	agg		_	tat	age	agt		_	tca	ava	agg		399
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cc2 30	c +-c	cc+	a+~	~~~	00+	-50	_		+		-45		+	<i>a</i> 20	160
cca ac															160
Pro Th		Pro	val	GIN		:?ro	GIÀ	Asn	ser		тте	Pro	ser	Asp	
-4					-35	_		_	_	-30			<u>.</u> ,		2.2
ct: qc															208
Leu Al	a Ser	Ser	Ser		Ser	Thr	Leu	Tyr	_	Cys	Ser	Thr	Cys		
-25				-20					-15					-10	_
ctc ag															256
Leu Ar			-5				_	1		_	_	5			
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Pro Hi	s His	Val	Ser	Arg	Ile	Ser	Trp	Thr	Leu	Ser	Val	Ser	Ser	His	

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286

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288

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Met Gly Ile Met Leu Leu Pro Arg Glu Cys Trp Lys Val Lys Asp Ser
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Lys Lys Tyr Lys Ser Cys Arg Glu Ser Val Leu Pro Ala Gln Ala Cys
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Thr Gly Glu Ser Pro Val Leu Ser Gly Val Arg Val Leu Gly Ile Arg
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Gln Lys Val Cys Tyr Leu Gly Ala Pro Cys Phe Gly Lys Arg Leu Ser
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Cys Leu Trp Ser Arg His Phe Gly Arg Leu Arg Arg Ala Ala Pro Leu	
1 5 10	
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291

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ctc ttt Leu Phe	e Ser	Leu :	Ile	Arg -15	Ser	His	Leu	Ser	Ile	ttg Leu	gct Ala	Phe	Val	Ala -5	281
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Glu Asr -55 acg cgg Thr Arg	y agg y Arg y aac y Asn	Ctc a Leu : agc a Ser : -20 gtg a	Sly atc Ile -35 atc Ile anc	Leu -50 ctt Leu ctg Leu	Glu gtt Val ggc Gly	gag Glu ggg Gly cag Gln tgc	Asn aga Arg aga Arg -15 acc	gct Ala aca Thr -30 cgg Arg	cag Gln -45 999 Gly ttc Phe	tcc Ser gcc Ala ttc Phe agc Ser	cgg Arg 999 Gly tcc Ser	aag Lys agg Arg -10	agc Ser -25 ctg Leu	tcc Ser -40 gcc Ala ggg Gly	161 209
Glu Asr -55 acg cgg Thr Arg act ggg Thr Gly gcc acg Ala Thr aag tgg Lys Cys 10	y agg y arg y aac y Asn y tct Ser -5 c cac s His	Ctc a Leu : agc a Ser : -20 gtg a Val 2 Val 0	atc Ile -35 atc Ile anc Xaa	Leu -50 ctt Leu ctg Leu agg Arg gtc Val	gtt Val ggc Gly gcc Ala gtr Val	gag Glu ggg Gly cag Gln tgc Cys 1 gnd Xaa	Asn aga Arg aga Arg -15 acc Thr ctm Leu	gct Ala aca Thr -30 cgg Arg acg Thr	cag Gln -45 999 Gly ttc Phe grh Xaa cat His 20	tcc Ser gcc Ala ttc Phe agc Ser 5 vwk Xaa	cgg Arg ggg Gly tcc Ser cgc Arg can	aag Lys agg Arg -10 agg Arg nmn Xaa	Glu agc Ser -25 ctg Leu tgg Trp ggg Gly	tcc Ser -40 gcc Ala ggg Gly gac Asp	161 209 257
Glu Asr -55 acg cgg Thr Arc act ggg Thr Gly gcc acc Ala Thr aag tgc Lys Cys	y agg y arg y aac y Asn y tct Ser -5 c cac s His	ctc a Leu : agc a Ser : -20 gtg a Val 2 gtg G Cag a Gln :	atc Ile -35 atc Ile anc xaa gaa Glu	Leu -50 ctt Leu ctg Leu agg Arg gtc Val 15 ctg	gtt Val ggc Gly gcc Ala gtr Val	gag Glu ggg Gly cag Gln tgc Cys 1 gnd Xaa	Asn aga Arg aga Arg -15 acc Thr ctm Leu	gct Ala aca Thr -30 cgg Arg acg Thr gga Gly	cag Gln -45 999 Gly ttc Phe grh Xaa cat His 20 gag	tcc Ser gcc Ala ttc Phe agc Ser 5 vwk Xaa	cgg Arg ggg Gly tcc Ser cgc Arg can Xaa	Gln aag Lys agg Arg -10 agg Arg nmn Xaa	Glu agc Ser -25 ctg Leu tgg Trp ggg Gly	tcc Ser -40 gcc Ala ggg Gly gac Asp aag Lys	161 209 257 305

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gag aag gga acc aag ccg cct tca gtt gag gat ggc ttc cag acc gtc Glu Lys Gly Thr Lys Pro Pro Ser Val Glu Asp Gly Phe Gln Thr Val -75 -65 -60	221
cct ctc atc act ccc ttg gag gtt aat cac tta cag ctg cct gct cca Pro Leu Ile Thr Pro Leu Glu Val Asn His Leu Gln Leu Pro Ala Pro -55 -50 -45	269
gaa aag gtg att gtg aag aca aga acg gaa tat cag ccg gaa cag aag Glu Lys Val Ile Val Lys Thr Arg Thr Glu Tyr Gln Pro Glu Gln Lys -40 -35 -30	317
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atc ctt gtc agc ctg gcc cta gct ttc ctt gcg tgc atc gtg ttc ctg  Ile Leu Val Ser Leu Ala Leu Ala Phe Leu Ala Cys Ile Val Phe Leu -10 -5 1 5	413
gtg gtt tac aaa gcc ttc acc tat gat cac agc tgc cca gag gat tcg Val Val Tyr Lys Ala Phe Thr Tyr Asp His Ser Cys Pro Glu Asp Ser 10 15 20	461
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aca gcc gtg ctt cca ctt gtg tct cac cag caa aac cat ctg ggt gga
                                                                       102
Thr Ala Val Leu Pro Leu Val Ser His Gln Gln Asn His Leu Gly Gly
agg ttt gca tct ctg gga tcc tca ggc att agg cac ggg
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                                                                       240
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tgttgtaaat gggattactt tttgcatttc tttchnnsaa ttgttcagtc agcatacagg
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aatgatactg atttttgt atg ttg att tta cat ctt gca act tta cta aat
                                                                       351
                    Met Leu Ile Leu His Leu Ala Thr Leu Leu Asn
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                                         -10
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Leu Phe Ile Ser Ser Asn Ser Phe
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aatattataa tgagg ccctggtatg accca aag agc agg gga Lys Ser Arg Gly agg agt ctg cgg Arg Ser Leu Arg -30 cca gat cac act Pro Asp His Thr -15 gtc ttc cct tct Val Phe Pro Ser 1 ccc atc tgc gtc Pro Ile Cys Val	ccc ckt gte Pro Xaa Val -45 gag tgg ccc Glu Trp Pro gta ctt gcc Val Leu Ala cag gtc acc Gln Val The 5 atc tct cac Ile Ser Gli	c cag act ct l Gln Thr Le t gat ctg tg c Asp Leu Cy -25 t ctg gtg tg a Leu Val Cy -10 c tgc aga ct r Cys Arg Le a ggt gcc tt n Gly Ala Ph	gggg cateu Gly His Gc tgc ttg gc tgc ttg gc tgc ttg gc cac agc ys His Ser cc cca agg eu Pro Arg 10 ct cac gat ne His Asp	atg gtc acc tca  Met Val Thr Ser  -50  gct ggc aac ctg  Ala Gly Asn Leu  -35  agg ctt ttt gtc  Arg Leu Phe Val  -20  gca tcc atc tct  Ala Ser Ile Ser  -5  aca ggg tca cat  Thr Gly Ser His  15  cct cac cca aat  Pro His Pro Asn	114 162 210 258 306

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1

- 5

302 aaa ggc agc tac gtm nnn tat gaa cct aca gaa ggt gag ccc agt gcc 302 Lys Gly Ser Tyr Val Xaa Tyr Glu Pro Thr Glu Gly Glu Pro Ser Ala 15 atc gtc cag atg gag adw nnc ttg gcc aag ggc agc gag 341 Ile Val Gln Met Glu Xaa Xaa Leu Ala Lys Gly Ser Glu 30 <210> 562 <211> 484 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 334..483 <221> sig peptide <222> 334..387 <223> Von Heijne matrix score 4.19999980926514 seq LIYLVSSFLALNQ/AS <400> 562 gttagttggg cagggctgaa gtgtatgtgg tgaggaaaag aggctcctac tgtagacagc 60 cttgttctac agatcctccc agaaatctct gggccaggtg gaacccaggg tcagagaggg 120 atgggagaga ggtttaattt tccatgataa ataaaaatct ataaaataat aaacaagaga 180 aaagagattg gaaacagcca ggttggagca gtgagtgagt aaggaaacct ggctgccctc 240 tccaqattcc ccaqqctctc aqaqaaqatc aqcaqaaaqt ctqcaaqass ctaaqaacca 300 teagecetea getgeacete eteceeteea agg atg aca aag geg sgv ete ate 354 Met Thr Lys Ala Xaa Leu Ile tat ttq qtc aqc aqc ttt ctt qcc cta aat caq qcc aqc ctc atc aqt 402 Tyr Leu Val Ser Ser Phe Leu Ala Leu Asn Gln Ala Ser Leu Ile Ser -10 -5 cgc tgt gac ttg gcc cag gtg ctg cag ctg gag gac ttg gat ggg ttt 450 Arg Cys Asp Leu Ala Gln Val Leu Gln Leu Glu Asp Leu Asp Gly Phe 10 gag ggt tac tcc ctg agt gac tgg ctg tgc tgg c 484 Glu Gly Tyr Ser Leu Ser Asp Trp Leu Cys Trp <210> 563 <211> 229 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 122..229 <221> sig peptide <222> 122..190 <223> Von Heijne matrix score 4.19999980926514 seq QLILLGIFRGIRH/QI <400> 563 60 gaaaggeete gaaggeageg teetaetega eeaccaagge aagacaagee aeetekattt agacggctaa gagagagga ggctgcttca aaatcaaatg aggtggtagc agtgcccaca 120 a atg gca cag tta ata atg tgg ctc aag aac cag tta ata ctc ttg ggg 169 Met Ala Gln Leu Ile Met Trp Leu Lys Asn Gln Leu Ile Leu Leu Gly ata ttt cgg gga ata aga cac cag att tat cta atc aga act ctt cag 217

303 Ile Phe Arg Gly Ile Arg His Gln Ile Tyr Leu Ile Arg Thr Leu Gln -5 atc agg caa tgg 229 Ile Arg Gln Trp <210> 564 <211> 352 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 80..352 <221> sig_peptide <222> 80..169 <223> Von Heijne matrix score 4.19999980926514 seq LAXTLSLTCSVSG/VS <400> 564 actttctgag agtcctggac ctcctgtgca agaacatgaa acatctgtgg ttcttccttc 60 teetggtgge aggteecag atg ggt eet gte eea ggt gea get gea gga gtm 112 Met Gly Pro Val Pro Gly Ala Ala Ala Gly Val -25 rgg ccc ayg amt ggc gaa ctt gcg grg acc ctg tcc ctc acc tgc agt 160 Xaa Pro Xaa Xaa Gly Glu Leu Ala Xaa Thr Leu Ser Leu Thr Cys Ser -15 -10 gto tot ggt gto too ato act agt tat tac tgg ago tgg ato ego car 208 Val Ser Gly Val Ser Ile Thr Ser Tyr Tyr Trp Ser Trp Ile Arg Gln gcc cca ggg aag ggg ccg gag tgg atc ggg cdk atc gat cat agc ggg 256 Ala Pro Gly Lys Gly Pro Glu Trp Ile Gly Xaa Ile Asp His Ser Gly 20 25 gat acc gac tac aat ccc tcc ctc cag agt cga gtc acc ctc tca gtg 304 Asp Thr Asp Tyr Asn Pro Ser Leu Gln Ser Arg Val Thr Leu Ser Val 30 35 40 gac acg tcg aag aac cag ttc tca ctg agg ttg ctt tct gtg agc gca 352 Asp Thr Ser Lys Asn Gln Phe Ser Leu Arg Leu Leu Ser Val Ser Ala 50 55 <210> 565 <211> 201 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 85..201 <221> sig peptide <222> 85..192 <223> Von Heijne matrix score 4.19999980926514 seq LPLFLCPLGMVET/SF <400> 565 agttetgege tgtgageegg ggeacaaaga geeetetgea etagegeege agaeegegga 60 ccagttggag gcatctgtcc accc atg tgg ttc cag aca cgt tca tgt ggc 111 Met Trp Phe Gln Thr Arg Ser Cys Gly -35 cac cat gac ccc gtc ggc atc aca ggg gta acc aag gtg atc ctc cct 159

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                                                                      113
                             Met Ser Tyr Val Val Thr Lys Thr Lys
                                              -105
geg att aat ggg aaa tac cat cgt ttc ttg ggt cgt cat ttc ccc cgc
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Ala Ile Asn Gly Lys Tyr His Arg Phe Leu Gly Arg His Phe Pro Arg
                    -95
                                        -90
ttc tat gtc ctg tac aca atc ttc atg aaa gga ttg cag atg tta tgg
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Phe Tyr Val Leu Tyr Thr Ile Phe Met Lys Gly Leu Gln Met Leu Trp
                                    -75
                -80
gct gat gcc aaa aag gct aga aga ata aag aca aat atg tgg aag cac
                                                                      257
Ala Asp Ala Lys Lys Ala Arg Arg Ile Lys Thr Asn Met Trp Lys His
            -65
                                -60
                                                     -55
aat ata aag ttt cat caa ctt cca tac cgg gag atg gag cat ttg aga
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Asn Ile Lys Phe His Gln Leu Pro Tyr Arg Glu Met Glu His Leu Arg
        -50
                            -45
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Gln Phe Arg Gln Asp Val Thr Lys Cys Leu Phe Leu Gly Ile Ile Ser
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                                             -25
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Ile Pro Pro Phe Ala Asn Tyr Leu Val Phe Leu Leu Met Tyr Leu Phe
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Pro Arg Gln Leu Leu Ile Arg
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score 4.19999980926514

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	104
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cca tgt caa gta acc tac aag ttc ttg ttt att ttg ctt gga cac gtc Pro Cys Gln Val Thr Tyr Lys Phe Leu Phe Ile Leu Leu Gly His Val 20 25 30	200
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g atg cag ggg ggc aac tcc ggg gtc cgc aag cgc gaa gag gag ggc gac Met Gln Gly Gly Asn Ser Gly Val Arg Lys Arg Glu Glu Glu Gly Asp	49 97
g atg cag ggg ggc aac tcc ggg gtc cgc aag cgc gaa gag gag ggc gac  Met Gln Gly Gly Asn Ser Gly Val Arg Lys Arg Glu Glu Glu Gly Asp  -75  -70  -65  ggg gct ggg gct gtg gct gcg ccg ccg gcc atc gac ttt ccc gcc gag Gly Ala Gly Ala Val Ala Ala Pro Pro Ala Ile Asp Phe Pro Ala Glu  -60  -55  -50	
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g atg cag ggg ggc aac tcc ggg gtc cgc aag cgc gaa gag gag ggc gac  Met Gln Gly Gly Asn Ser Gly Val Arg Lys Arg Glu Glu Glu Gly Asp  -75  -70  -65  ggg gct ggg gct gtg gct gcg ccg ccg gcc atc gac ttt ccc gcc gag Gly Ala Gly Ala Val Ala Ala Pro Pro Ala Ile Asp Phe Pro Ala Glu  -60  -55  -50  ggc ccg gac ccc gaa tat gac gaa tct gat gtt cca gca kaa atc cag Gly Pro Asp Pro Glu Tyr Asp Glu Ser Asp Val Pro Ala Xaa Ile Gln  -45  -40  -35  gtg tta aaa gaa ccc cta caa cag cca acc ttc cct ttt gca gtt gca Val Leu Lys Glu Pro Leu Gln Gln Pro Thr Phe Pro Phe Ala Val Ala  -30  -25  -20  aac caa ctc ttg ctg gtt tct ttg ctg gag cac ttg agc cac gtg cat Asn Gln Leu Leu Leu Val Ser Leu Leu Glu His Leu Ser His Val His	97 145
g atg cag ggg ggc aac tcc ggg gtc cgc aag cgc gaa gag gag ggc gac  Met Gln Gly Gly Asn Ser Gly Val Arg Lys Arg Glu Glu Glu Gly Asp  -75  -70  -65  ggg gct ggg gct gtg gct gcg ccg ccg gcc atc gac ttt ccc gcc gag Gly Ala Gly Ala Val Ala Ala Pro Pro Ala Ile Asp Phe Pro Ala Glu  -60  -55  -50  ggc ccg gac ccc gaa tat gac gaa tct gat gtt cca gca kaa atc cag Gly Pro Asp Pro Glu Tyr Asp Glu Ser Asp Val Pro Ala Xaa Ile Gln  -45  -40  -35  gtg tta aaa gaa ccc cta caa cag cca acc ttc cct ttt gca gtt gca Val Leu Lys Glu Pro Leu Gln Gln Pro Thr Phe Pro Phe Ala Val Ala  -30  -25  -20  aac caa ctc ttg ctg gtt tct ttg ctg gag cac ttg agc cac gtg cat Asn Gln Leu Leu Leu Val Ser Leu Leu Glu His Leu Ser His Val His  -15  -10  -5  -70  -65  -70  -65  -70  -65  -70  -70  -70  -70  -70  -70  -70  -7	97 145 193
g atg cag ggg ggc aac tcc ggg gtc cgc aag cgc gaa gag gag ggc gac  Met Gln Gly Gly Asn Ser Gly Val Arg Lys Arg Glu Glu Glu Gly Asp  -75  -70  -65  ggg gct ggg gct gtg gct gcg ccg ccg gcc atc gac ttt ccc gcc gag Gly Ala Gly Ala Val Ala Ala Pro Pro Ala Ile Asp Phe Pro Ala Glu  -60  -55  -50  ggc ccg gac ccc gaa tat gac gaa tct gat gtt cca gca kaa atc cag Gly Pro Asp Pro Glu Tyr Asp Glu Ser Asp Val Pro Ala Xaa Ile Gln  -45  -40  -35  gtg tta aaa gaa ccc cta caa cag cca acc ttc cct ttt gca gtt gca Val Leu Lys Glu Pro Leu Gln Gln Pro Thr Phe Pro Phe Ala Val Ala  -30  -25  -20  aac caa ctc ttg ctg gtt tct ttg ctg gag cac ttg agc cac gtg cat Asn Gln Leu Leu Leu Val Ser Leu Leu Glu His Leu Ser His Val His  -15  -10  -5  1  gaa cc	97 145 193
g atg cag ggg ggc aac tcc ggg gtc cgc aag cgc gaa gag gag ggc gac  Met Gln Gly Gly Asn Ser Gly Val Arg Lys Arg Glu Glu Glu Gly Asp  -75  -70  -65  ggg gct ggg gct gtg gct gcg ccg ccg gcc atc gac ttt ccc gcc gag Gly Ala Gly Ala Val Ala Ala Pro Pro Ala Ile Asp Phe Pro Ala Glu  -60  -50  ggc ccg gac ccc gaa tat gac gaa tct gat gtt cca gca kaa atc cag Gly Pro Asp Pro Glu Tyr Asp Glu Ser Asp Val Pro Ala Xaa Ile Gln  -45  -40  -35  gtg tta aaa gaa ccc cta caa cag cca acc ttc cct ttt gca gtt gca Val Leu Lys Glu Pro Leu Gln Gln Pro Thr Phe Pro Phe Ala Val Ala  -30  -25  aac caa ctc ttg ctg gtt tct ttg ctg gag cac ttg agc cac gtg cat Asn Gln Leu Leu Leu Val Ser Leu Leu Glu His Leu Ser His Val His  -15  -10  -5  Glu <a href="#">-5</a> 1  gaa cc Glu <a href="#">-20</a> 220  230  240  250  569 <a href="#">-210</a> -5  1  260  -5  1  27  28  28  29  20  21  20  21  21  21  21  21  21  21	97 145 193

306

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                                                                      110
                   Met Val Phe Tyr Cys Phe Ala Leu Cys Ile Ile
                           -15
ctt att tgt gtt atg tct tgt cgc cac ctg gg
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gtc tcc tgc aca gct ggc tct gcg tgt gct ctt tct cta ttg caa ttc
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Val Ser Cys Thr Ala Gly Ser Ala Cys Ala Leu Ser Leu Leu Gln Phe
                    -25
                                         -20
cct gtc ttg ata act cag ctc tgt cta ggc aaa ggg caa agt gaa ccc
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Pro Val Leu Ile Thr Gln Leu Cys Leu Gly Lys Gly Gln Ser Glu Pro
                                     -5
                -10
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                                                                      194
Ile Gly Pro Leu Gln Asp Phe Val Ser Leu Glu Ser Thr Ser His Phe
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Tyr Ser Phe Phe
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Met Thr Arg Arg Thr -20 -15	
tet etg tgg tge tge age eet tet tee aga aca tee age tee etg tee Ser Leu Trp Cys Cys Ser Pro Ser Ser Arg Thr Ser Ser Ser Leu Ser	341
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ctatggcttg tag atg gcc ttc tat ctc tgg tgt ttt cat gcg gtc ttt	169
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gcggccttcc gcagtgtttg tgtccctggg tacttgagat tagggagtgg tg atg act Met Thr	178
ctt aac gag cat gct gcc ttc aag cat ctg ttt aac aaa gca cat ctt	226
Leu Asn Glu His Ala Ala Phe Lys His Leu Phe Asn Lys Ala His Leu	·
-30 -25 -20	

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Arg Pro Leu Gln Pro Phe Leu Glu His Leu Ala Gly Ser Gly Ile Thr

PCT/IB99/00712

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Leu Phe Glu Arg Ala Glu Ser Leu Tyr Thr Leu Ile Ser Lys Phe Xaa -25 -20 -15	
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Met	
ttt gtg aat aga acc tgt ttt aat tct tcc ttt cca atc tgg atg cct	165
Phe Val Asn Arg Thr Cys Phe Asn Ser Ser Phe Pro Ile Trp Met Pro	
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Met Xaa Gly Ser Ser Arg Xaa Xaa Gly -35 -30	51

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311	
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ttt ctg skc gcc gtc agc gcc agt agc tcg cma gca Phe Leu Xaa Ala Val Ser Ala Ser Ser Ser Xaa Ala	
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ggt ttt cat ggt ttt tct aaa tac aca gtt tca cgt Gly Phe His Gly Phe Ser Lys Tyr Thr Val Ser Arg 1 5 10	
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score 4.09999990463257

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314

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Met Lys Ser Asn Leu Thr Leu Leu Thr Cys Leu Xaa Leu Xaa Gly

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317

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Glu Ile Leu Leu Leu Ile Thr Ile Ile Tyr Ser Tyr Leu Glu Ser
                -10
                                    -5
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Leu Val Lys Phe Phe Ile Pro Gln Arg Arg Lys Ser Val Ala Gly Glu
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                                                                      120
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gca gcc ctc gtg aca ttt gga agc att ttt gga tat aag cdg aga ggt
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                        -25
                                             -20
ggt gtt eeg tet ttg att get ggt ett ttt gtd gga tgt ttg gee gge
                                                                      268
Gly Val Pro Ser Leu Ile Ala Gly Leu Phe Val Gly Cys Leu Ala Gly
                    -10
tat nsa gct tac cgt gtc tcc aat gac aaa cga gat gta a
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Tyr Xaa Ala Tyr Arg Val Ser Asn Asp Lys Arg Asp Val
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•						-		-15	5				-10	)	Ala	
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ctg t Leu T 10																147
agt c Ser H																195
atg g Met V																219
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ttt a Phe T			_										tgt	ggc		164
gga a Gly S 1	er															212
gjà aaa																215
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320

Gln Pro Leu Pro Pro Gly Phe Lys Pro Xaa Ser Cys Leu Ser Leu Leu -10 agt aay tsa gat tac agg cat gca cca cca ttc ctg gct aat ttt kgw 380 Ser Asn Xaa Asp Tyr Arg His Ala Pro Pro Phe Leu Ala Asn Phe Xaa att ttt cat aga gat gga gtt tca cca 407 Ile Phe His Arg Asp Gly Val Ser Pro <210> 597 <211> 274 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 90..272 <221> sig_peptide <222> 90..254 <223> Von Heijne matrix score 4 seq LHQGLCLPQRVHC/SL getgacegrg egeacseege eeceggsgee atetteeega eegegageeg teeaggtete 60 agtgctrtgc ccccccaga gcctagagg atg ttt cat ggg atc cca gcc acg 113 Met Phe His Gly Ile Pro Ala Thr -55 -50 ccg ggc ata gga gcc cct ggg aac aag ccg gag ctg tat gag gta cga 161 Pro Gly Ile Gly Ala Pro Gly Asn Lys Pro Glu Leu Tyr Glu Val Arg -40 -35 caa cat ggc aga gct gtt tgc ggt ggt gaa gac aat gca agc cct gga 209 Gln His Gly Arg Ala Val Cys Gly Glu Asp Asn Ala Ser Pro Gly -25 -20 gaa ggc cta cat caa gga ctg tgt ctc ccc cag cga gta cac tgc agc 257 Glu Gly Leu His Gln Gly Leu Cys Leu Pro Gln Arg Val His Cys Ser -10 -5 274 ctq ctc ccq gct cct gg Leu Leu Pro Ala Pro <210> 598 <211> 417 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 343..417 <221> sig_peptide <222> 343..408 <223> Von Heijne matrix sed IFLSVLNFIFTLS/FS gcatctaqaa qtacaaqttq atgattattq tccatttgat agagacactg gaagggtgtc 60 agtgtaaaca ctggccatgt gaagattgag cctgttgatg gtttcttttg tatcatagga tgccacgtca ccaactaggg aattctgccc aatcagttga gccaaatagt gctgtcctat tgtaaaattg tttaatctgt gtgcttgtgt gtgtgcttgt cagaatttgt gaatcataga 240 attgttttaa ctggaagaag acccccaaga tcatctgctt caaccccttc cttcctctct 300

321	
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ttt gta agt tgt aac cta ttt ttg tct gtr ttg aat ttc ctt ttt ttg Phe Val Ser Cys Asn Leu Phe Leu Ser Val Leu Asn Phe Leu Phe Leu -15 -10 -5	402
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Ser Glu Ala Leu Pro Ser Leu Ala Gly Asp Pro Val Ala Val Glu Ala -20 -15 -10	225
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322
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                                                                       120
aggaacattc taattcaaat tcatatagtt ggtccttg atg cgg ttg gcg atg gtg
                                                                       176
                                           Met Arg Leu Ala Met Val
caa ttg gtg ctc aac aat ttg aag act ttt tat ccc ttc gca gat cat
                                                                       224
Gln Leu Val Leu Asn Asn Leu Lys Thr Phe Tyr Pro Phe Ala Asp His
                    -25
                                         -20
gat ctt gca gag ctt cca gtt agt tca cct ctt tgt cat gcg gtt cta
                                                                       272
Asp Leu Ala Glu Leu Pro Val Ser Ser Pro Leu Cys His Ala Val Leu
                -10
                                     -5
aaa act ctt caa tgt tgg gaa caa gtt ctt ctc cga cga ctt gaa atc
                                                                       320
Lys Thr Leu Gln Cys Trp Glu Gln Val Leu Leu Arg Arg Leu Glu Ile
                            10
cat ggt ggg cca cct caa aat tat atc gca agt cat acc gcc gan nag
                                                                       368
His Gly Gly Pro Pro Gln Asn Tyr Ile Ala Ser His Thr Ala Xaa Xaa
    20
agt ttg tct gca ggt cct gca att ctt cgc cac aaa gct tta ctg gaa
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Ser Leu Ser Ala Gly Pro Ala Ile Leu Arg His Lys Ala Leu Leu Glu
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cct a
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Pro
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324

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-25 -20 -15 cgg gag ttg cgc tac ctg agc gcg gcc acc ggc cac cct atc gcg aca 1	155
Arg Glu Leu Arg Tyr Leu Ser Ala Ala Thr Gly His Pro Ile Ala Thr -10 -5 1 5	
ccg cgg cct atc ggt acc ntt gtg aag gct ttc cgt gca cat cgg gtc Pro Arg Pro Ile Gly Thr Xaa Val Lys Ala Phe Arg Ala His Arg Val	203
10 15 20	
acc agt gaa aag ttg tgc aga gcc caa cat gag ctt cat ttc caa gct Thr Ser Glu Lys Leu Cys Arg Ala Gln His Glu Leu His Phe Gln Ala	251
. 25 30 35	
gcc acc tat ctc tgc ctc ctg cgt asa tcc gga aac atg tgg ccc tac Ala Thr Tyr Leu Cys Leu Leu Arg Xaa Ser Gly Asn Met Trp Pro Tyr	299
40 45 50	
atc agg aat ttc atg gca agg gtg agc gct cgg tgg agg agt ctg ctg  Ile Arg Asn Phe Met Ala Arg Val Ser Ala Arg Trp Arg Ser Leu Leu	347
55 60 65	
	395
Ala Trp Trp Val Ser Ser Cys Pro Ile Ser Leu Glu Gly Arg Ala Gly 70 75 80 85	
	426
Ser His Glu His Gly Glu Tyr Pro Trp Met 90 95	
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Met Ile Thr Asp Val Gln Leu Ala Ile Phe Ala Asn Met	
-25 -20 ctg ggc gtg tcg ctc ttc ttg ctt gtc gtt ctc tat cac tac gcg gcc 1	157
Leu Gly Val Ser Leu Phe Leu Leu Val Val Leu Tyr His Tyr Ala Ala	13,
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gct ttg agc ttc ctt tgc tct tta tcg caa aat gca ttg aat att tcc Ala Leu Ser Phe Leu Cys Ser Leu Ser Gln Asn Ala Leu Asn Ile Ser -10 -5 1	342
ctc att tct cgt aag Leu Ile Ser Arg Lys 5	357
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act cag tta aga aat tct tcc tta gcc atg Thr Gln Leu Arg Asn Ser Ser Leu Ala Met 1 5	201
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327	
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act gtc ggg tgc gct aca gcc agc tcc tgg ggc tgy acg agc agg gg Thr Val Gly Cys Ala Thr Ala Ser Ser Trp Gly Cys Thr Ser Arg -10 -5 1	281
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cgg cag aag cag ctg gcc gcc tgg tgc tcg ctg gtc ctg tcc ttc tgc Arg Gln Lys Gln Leu Ala Ala Trp Cys Ser Leu Val Leu Ser Phe Cys -15 -10 -5	150
cgc ctg cac aaa cag tcc agc atg acg gtg atg gaa gct cag gag agc Arg Leu His Lys Gln Ser Ser Met Thr Val Met Glu Ala Gln Glu Ser 1 5 10	198
ccg ctc ttc aac aac gtc aag cta cag cga aag ctt cct gtg g Pro Leu Phe Asn Asn Val Lys Leu Gln Arg Lys Leu Pro Val 15 20 25	241
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Met Arg Leu His gta cat tcc ctt tct ccc ttt tcc ttt gct tgt ctc cct ttt ctg tcc Val His Ser Leu Ser Pro Phe Ser Phe Ala Cys Leu Pro Phe Leu Ser	165

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WO 99/53051 PCT/	IB99/00712
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ttg gaa aga atg tgc att ctg caa ttg ttg agt gct gtg ttg tat aga Leu Glu Arg Met Cys Ile Leu Gln Leu Ser Ala Val Leu Tyr Arg -15 -10 -5 1 ttt g Phe	338
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ttattatact tcttaactaa tcaactatwm cyttacccat ctagccaaag tagactaccc
                                                                      180
atat atg ttt ctt gac cat gtc agg ttt tta acc tcc ata tct ttt ctt
                                                                      229
    Met Phe Leu Asp His Val Arg Phe Leu Thr Ser Ile Ser Phe Leu
                         -15
gct ctg gtc ctg tgg aat gtc ttt ctc aac tct acc cgt ctg g
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Ala Leu Val Leu Trp Asn Val Phe Leu Asn Ser Thr Arg Leu
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                                                                      101
Pro Ala Leu Leu Thr Ser Ser Glu Leu Pro Ala Leu Ala Ser Gln Ile
cat tgc cgc gtc c
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His Cys Arg Val
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331

331	
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tgc tcg gga ccg ctt tcc ctc cgt tcc cct cgg ctt ccc cct ctc ttt Cys Ser Gly Pro Leu Ser Leu Arg Ser Pro Arg Leu Pro Pro Leu Phe -25 -20 -15	162
tgc act ttt ctt tcc ctt tct ttg cat ccc tgg ggg ggt ttc ttt ttg Cys Thr Phe Leu Ser Leu Ser Leu His Pro Trp Gly Gly Phe Phe Leu -10 -5 1	210
tgt gcc tgg att tct bkt ttc ctc ccg tgg gtg tgt gtg tgk gcg gg Cys Ala Trp Ile Ser Xaa Phe Leu Pro Trp Val Cys Val Xaa Ala 5 10 15	257
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tca aag act agg acc aat gat gga aaa att aca tat ccg cct ggg gtc Ser Lys Thr Arg Thr Asn Asp Gly Lys Ile Thr Tyr Pro Pro Gly Val -85 -80 -75	103
aag gaa ata tca gat aaa ata tct aaa gag gag atg gtg aga cga tta Lys Glu Ile Ser Asp Lys Ile Ser Lys Glu Glu Met Val Arg Arg Leu -70 -65 -60 -55	151
aag atg gtt gtg aaa act ttt atg gat atg gac cag gac tct gaa gaa Lys Met Val Val Lys Thr Phe Met Asp Met Asp Gln Asp Ser Glu Glu -50 -45 -40	199
gaa aag gag ctt tat tta aac cta gct tta cat ctt gct tca gat ttt Glu Lys Glu Leu Tyr Leu Asn Leu Ala Leu His Leu Ala Ser Asp Phe -35 -30 -25	247
ttt ctc aag cat cct gat aaa gat gtt cgc tta ctg gta gcc tgc tgc Phe Leu Lys His Pro Asp Lys Asp Val Arg Leu Leu Val Ala Cys Cys -20 -15 -10	295
ctt gct gat att ttc agg att tat gct cct gaa gct cct tac aca tcc Leu Ala Asp Ile Phe Arg Ile Tyr Ala Pro Glu Ala Pro Tyr Thr Ser -5 1 5 10	343
cct aag gg Pro Lys	351
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201	•

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332

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aga ct Arg Le	u Phe	Gly -20	Cys	Phe	Pro	Ser	Asp -15	Leu							403
tct ag Ser Se								ca							432
<210><211><211><212><213>	233 DNA	sapie	ens												
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tac ct Tyr Le								agt	act				atg	aac	162
cgt ct Arg Le 5															210
tgc ct	a ccc u Pro					ca									233
				25											5,
Cys Le 20 <210> <211> <212> <213>	380 DNA	sapie	ens	25											in the second se
20 <210> <211> <212>	380 DNA Homo s		ens	25									·		` <u>`</u>
20 <210> <211> <212> <213> <220> <221>	380 DNA Homo s CDS 1037 sig_pe	78 eptic 7 eijne	le = mat	:rix	36743										` <u>`</u>

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ggt g Gly G			cag	ggc				aag	gcg				ggc	agt		99
cgg a Arg I 15	le '	Thr	Ala	Ala	Val 20	Ile	Glu	His	Leu	Glu 25	Arg	Leu	Ala	Leu	Val 30	147
gac t Asp P																195
ttc g Phe A	la i	gac Asp	cgg Arg 50	cta Leu	cgc Arg	gcc Ala	gtg Val	gac Asp 55	aca Thr	gac Asp	Gly 999	gtg Val	gag Glu 60	ccc Pro	atg Met	243
gaa t Glu S	er '															291
gta g Val G 8																339
gtg g Val G 95													tt			380
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ccc g Pro G	ly (	gaa Glu -70	cgt Arg	ctg Leu	tgt Cys	aac Asn	ttg Leu -65	gag Glu	gag Glu	ggc Gly	agc Ser	ccg Pro -60	ggc Gly	agc Ser	ggc Gly	101
acc t Thr T																149
atg a Met L -40	ag :	agc Ser	agc Ser	gag Glu	aat Asn -35	ggc Gly	gcg Ala	ctt Leu	cca Pro	gtg Val -30	gtg Val	tct Ser	gta Val	gtg Val	aga Arg -25	197
gaa a Glu T	ca ( Chr (	gag Glu	pad Ser	cag Gln -20	t.ta Leu	ctg Leu	cca Pro	gat Asp	gtg Val -15	gga Gly	gct Ala	att. Ile	gta Val	acc Thr -10	tgr Cys	245
aag t Lys S										a					•	276

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336

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gtg agt gtt aca gtt ctt aaa gat ggt gtg gct gga gtt tgt ttc ttc Val Ser Val Thr Val Leu Lys Asp Gly Val Ala Gly Val Cys Phe Phe -20 -15 -10	282
aga cgt tca gat gcg tct gaa gtt tct tcc ttc tgg Arg Arg Ser Asp Ala Ser Glu Val Ser Ser Phe Trp -5 5	318
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ctc atg atc agt aat act gag ctt ttt ttc ata cgc ttc ttg act gca Leu Met Ile Ser Asn Thr Glu Leu Phe Phe Ile Arg Phe Leu Thr Ala -35 -30 -25 -20	100
tgt atg cct tct ttt gaa aag tgt ctg ttc tta tct ttt gcc cac ttc Cys Met Pro Ser Phe Glu Lys Cys Leu Phe Leu Ser Phe Ala His Phe -15 -10 -5	148
ttg atg gga aga acc cac cgt g Leu Met Gly Arg Thr His Arg 1	170
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agt ctc tta tca gat ata tta ttt gca aat att ttc tcc cat tct tgg Ser Leu Leu Ser Asp Ile Leu Phe Ala Asn Ile Phe Ser His Ser Trp	161

337 -10 -5 1 gac gtc ttt cca ctt tct ttt ctt ttc ttt tct tt 196 Asp Val Phe Pro Leu Ser Phe Leu Phe Phe Ser <210> 631 <211> 339 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 53..337 <221> sig_peptide <222> 53..304 <223> Von Heijne matrix score 3.90000009536743 seg SSLLIILLPNTQD/LR <400> 631 agttccgacg aaaaatggcg gggtctcctg agttggtggt ccttgaccct cc atg gga 58 Met Gly caa gga gct cgc ggc tgg cac aga gag cca ggc ctt ggt ctc cgc cac 106 Gln Gly Ala Arg Gly Trp His Arg Glu Pro Gly Leu Gly Leu Arg His -80 -75 tee eeg aga aga ett teg ggt geg etg eac ete gaa geg gge tgt gae 154 Ser Pro Arg Arg Leu Ser Gly Ala Leu His Leu Glu Ala Gly Cys Asp -60 -55 cga aat gct aca act gtg cgg ccg ctt cgt gca aaa shc ggg gac gct 202 Arg Asn Ala Thr Thr Val Arg Pro Leu Arg Ala Lys Xaa Gly Asp Ala -40 -45 ctg ccg gag gag att cgg gag ccc gct ctg cga gat gcg cag tgg gta 250 Leu Pro Glu Glu Ile Arg Glu Pro Ala Leu Arg Asp Ala Gln Trp Val -30 -25 cgg gac cag tta gcc agt tct tta ctc atc atc ctc tta ccc aac acc 298 Arg Asp Gln Leu Ala Ser Ser Leu Leu Ile Leu Leu Pro Asn Thr -10 -15 cag gat ctt agg att cag aaa gat ccc aca cca ggc ccg gg 339 Gln Asp Leu Arg Ile Gln Lys Asp Pro Thr Pro Gly Pro <210> 632 <211> 433 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 171..431 <221> sig_peptide <222> 171..314 <223> Von Heijne matrix score 3.79999995231628 seq NSLLLLLCLYIYP/HS

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aaq aga aaq atc agt qtq tqt caa caa act tqq qcc tta tta tqc aaq	224											
Lys Arg Lys Ile Ser Val Cys Gln Gln Thr Trp Ala Leu Leu Cys Lys -45 -40 -35	224											
aac, ttt ctt aaa aaa tgg aga atg aaa aga gag tcc tta atg gaa tgg	272											
Asn Phe Leu Lys Lys Trp Arg Met Lys Arg Glu Ser Leu Met Glu Trp -30 -25 -20 -15												
ctg aat tca ttg ctc cta cta ctt tgt ttg tat ata tat cct cat agt	320											
Leu Asn Ser Leu Leu Leu Leu Cys Leu Tyr Ile Tyr Pro His Ser -10 -5 1												
cat caa gta aat gaw tdd tct tca ctg ctt acc atg gac ctg gga cgg	368											
His Gln Val Asn Xaa Xaa Ser Ser Leu Leu Thr Met Asp Leu Gly Arg  5 10 15												
gta gat rnn tkt aat gaa tcc aga ttt tct gtt gta tac aca cct gtc	416											
Val Asp Xaa Xaa Asn Glu Ser Arg Phe Ser Val Val Tyr Thr Pro Val 20 25 30												
acc aac acg acc cct gg	433											
Thr Asn Thr Thr Pro 35												
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Met -30												
$\cdot$	304											
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Cys Thr Cys Leu Cys Val Cys Leu Tyr Met Tyr Asn Met Gln Phe Leu												
-25 -20 -15	150											
kyt ttt gtg ttt gtk tgc gww ttg cta aag tgt atg agt gtg cct ttg	152											
Xaa Phe Val Phe Val Cys Xaa Leu Leu Lys Cys Met Ser Val Pro Leu												
-10 -5 1 tg	154											
010 624												
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15

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<211> 172

<212> DNA

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     ccc aga cag aga aac tca atg tgt ttg ctt tta gac gtc agc tct rcc
                                                                          103
     Pro Arg Gln Arg Asn Ser Met Cys Leu Leu Leu Asp Val Ser Ser Xaa
                 -15
                                     -10
     aaq aqc aca qat aat tth cya rtc gww wtt ttg att att tat ctg
                                                                          151
     Lys Ser Thr Asp Asn Xaa Xaa Xaa Xaa Leu Ile Ile Tyr Tyr Leu
     att acc aga aaa ggg cca ggg
                                                                          172
     Ile Thr Arg Lys Gly Pro Gly
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     ageagaeece teaetetgaa geeeceggat eeaageagg atg age tge caa met
                                                                          114
                                                Met Ser Cys Gln Xaa
     mag ctt gct cdg acc ttg act tgg ctc atg atc cgt gga aga cat ccc
                                                                          162
     Xaa Leu Ala Xaa Thr Leu Thr Trp Leu Met Ile Arg Gly Arg His Pro
                 -35
                                     -30
                                                         -25
     tac ctg acc cgt cga tca gcc cga aac ttc aac atc ttt ttg gca gct.
                                                                          210
     Tyr Leu Thr Arg Arg Ser Ala Arg Asn Phe Asn Ile Phe Leu Ala Ala
             -20
                                 -15
     ccg tcc cca gtt tgg cag cct cag agg acc cgc cga ccc cag k
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WO 99/53051 343 taagagetga gegeastgac aactagggge eggacegteg eaggaggegt eegetggata cettececet teeetgacet agagetetae agetgetgee teggtactga cegagggtte 240 ccagagetgt ctyaccattg caaaaacgtt atagcaacag cetetgatta egac atg 297 get gag ate ace aat ate ega eet age tit gat gig tea eeg gig gig 345 Ala Glu Ile Thr Asn Ile Arg Pro Ser Phe Asp Val Ser Pro Val Val -35 -30 gee gge etc atc ggg gee tet gtg etg gtg tgt gte teg gtg acc 393 Ala Gly Leu Ile Gly Ala Ser Val Leu Val Val Cys Val Ser Val Thr -15 -10 gtc ttt gtc tgg tca tgc tgc crc cag cag gca gag aag aag cac aag 441 Val Phe Val Trp Ser Cys Cys Xaa Gln Gln Ala Glu Lys Lys His Lys aac cca cca tac aag ttt att cac atg ctc aaa ggc wtc agc 483 Asn Pro Pro Tyr Lys Phe Ile His Met Leu Lys Gly Xaa Ser <210> 642 <211> 309 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 235..309 <221> sig_peptide <222> 235..279 <223> Von Heijne matrix score 3.79999995231628 seq ILTMLILLIHEHG/IF <400> 642 attratetat gtgtetgttg ttataegaat ateatgetgt tttggtttet atateettgt 60 aatatgtttt gaagtcaggt agtgtgatgc ctccagattt gttctttttg gtcaggattg 120 ctttggctgw tttgggttcw wttwtggttc catacaaatt ttaggattat tttttctatg 180 tctgtgaaaa gtggcatggg tattacattc aatctgtaga ttgctttgga tagt atg 237 Met gtc att tta act atg tta att ctt tta atc cat gag cat ggt att ttc 285 Val Ile Leu Thr Met Leu Ile Leu Leu Ile His Glu His Gly Ile Phe -10 309 ttt tca ctt gtt tgt gtc ctc ttc Phe Ser Leu Val Cys Val Leu Phe <210> 643 <211> 245 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 147..245 <221> sig_peptide <222> 147..233 <223> Von Heijne matrix

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<221> misc feature

<222> 61

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-25 tac aca cca cag cac agc ccg ctc aca cac aca cac aca tgc acc cca	221
Tyr Thr Pro Gln His Ser Pro Leu Thr His Thr His Thr Cys Thr Pro -20 -15 -10 -5	221
ccc agc aca gct cac cca cgc ggg Pro Ser Thr Ala His Pro Arg Gly 1	245
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gca gtt tcg aat ttt aat aaa ctt tta tgg gga gva ag Ala Val Ser Asn Phe Asn Lys Leu Leu Trp Gly Xaa -5 5	211
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ctt gct tgc ttc tct ctc ttt ggc wtt ctt cct cag ggg ctc ctt atc Leu Ala Cys Phe Ser Leu Phe Gly Xaa Leu Pro Gln Gly Leu Leu Ile -10 -5 1	98

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 ccccggagtt cagaggtcta aggaagagga gataaatata tgaaggtgct gtttggcaca
                                                                        120
 gaatttaata gggaagaaag agacagtata actcaccagt gctgggtctc atcatcctgc
                                                                        180
 aatttcdgaa caactatgaa tacaaaaaga attttaaaaat cccagtcctg cctagaaagg
                                                                        240
 ggaagtcatc tctaaat atg gtg gcc ctg ggg cag ctg gcc tdc ctg cca
                                                                        290
                    Met Val Ala Leu Gly Gln Leu Ala Xaa Leu Pro
                                     -15
 gge nbc tdc cat ggg ggc ctt tct gca gtg act gtg gtt ctt ccc att
                                                                        338
 Gly Xaa Xaa His Gly Gly Leu Ser Ala Val Thr Val Val Leu Pro Ile
                                                                        347
 tta ctc tgt
 Leu Leu Cys
     10
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 gttgagcaat tittcat atg ccc gtt tca tit gtc tgt ctt ctt ttc aga
                                                                        110
                     Met Pro Val Ser Phe Val Cys Leu Leu Phe Arg
                     -15
                                         -10
 aat gtt tat tca aat cta ttg cct tct ttt ttt
                                                                        143
 Asn Val Tyr Ser Asn Leu Leu Pro Ser Phe Phe
                  1
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<211> 232

<212> DNA

346

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								gga Gly							gct Ala	400
	_	_	ctc Leu	_	_	_										419
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atc Ile	ttg Leu -70	aat Asn	ggc Gly	ttc Phe	cgg Arg	ggc Gly -65	cat His	gcc Ala	aca Thr	gat Asp	tcc Ser -60	gtg Val	aag Lys	aac	tcc	105
atg Met -55	gaa Glu	agc Ser	atg Met	aac Asn	act Thr -50	gac Asp	atg Met	gtg Val	atc Ile	atc Ile -45	cca Pro	gly aaa	ggt Gly	ctg Leu	acc Thr -40	153
tca Ser	cag Gln	ctt Leu	cag Gln	gtg Val -35	ctg [.] Leu	gat Asp	gtc Val	gtg Val	gtc Val -30	tac Tyr	aag Lys	cca Pro	ctg Leu	aat Asn -25	gac Asp	201
agt Ser	gtg Val	cgg Arg	gcc Ala -20	cag Gln	tac Tyr	tcc Ser	aac Asn	tgg Trp -15	ctt Leu	ctg Leu	gct Ala	Gly 999	aac Asn -10	ctg Leu	gcg Ala	249
ctg Leu	agc Ser	cca Pro -5	acc Thr	Gly 999	aat Asn	gct Ala	aag Lys 1	aag Lys	cca Pro	ccc Pro	ctg Leu 5	ggc Gly	ctc Leu	ttt Phe	ctg Leu	297
Glu 10	Trp	Val	Met	Val	Ala 15	Trp	Asn	agc Ser	Ile	Ser 20	Ser	Glu	Ser	Ile	Val 25	345
caa Gln	Gly 999	whc Xaa	aaa Lys	gaa Glu 30	gtg Val	cca Pro	tat Tyr	ctc Leu	crg Xaa 35	caa Gln	ctt Leu	gga Gly	gga Gly	gga Gly 40	aga Arg	393
cga																396

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<222> 67..168
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                                                                        60
ggatte atg ate tgt ace act gtt tat att ace atg get eet tae tgt
                                                                       108
       Met Ile Cys Thr Thr Val Tyr Ile Thr Met Ala Pro Tyr Cys
                           -20
cta tca aac tgt tta ctt thw caw agt tgg ggc ctg cat ttg tat aga
                                                                       156
Leu Ser Asn Cys Leu Leu Xaa Xaa Ser Trp Gly Leu His Leu Tyr Arg
ttt cta gcc ccc at
                                                                       170
Phe Leu Ala Pro
<210> 653
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ctctgggaac aggaaagtca ggaaccctgc ctttcaggaa ctgctgtatc tcagtcggct
                                                                       120
tottcatttc atg gtt tot ctc tgt gta gct gct tta ttt cct ctt cag
                                                                       169
           Met Val Ser Leu Cys Val Ala Ala Leu Phe Pro Leu Gln
gct tac ggg
                                                                       178
Ala Tyr Gly
    1
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349

<222> 137..196

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atc cta tat ata tac cac ctg cgg g  Ile Leu Tyr Ile Tyr His Leu Arg  1 5	197
<210> 657 <211> 246 <212> DNA <213> Homo sapiens	
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ttk gtt ttt kgt ttt tkg ttt ttg ara cgg asy ttg cyc tkg ycg ccc Xaa Val Phe Xaa Phe Xaa Phe Leu Xaa Arg Xaa Leu Xaa Xaa Pro -10 -5 1	159
agg ctg gag tgc aat ggm aar ayc tcg gcy cac tgm aac ctc cgc ctc Arg Leu Glu Cys Asn Gly Lys Xaa Ser Ala His Xaa Asn Leu Arg Leu 5 10 15	207
ctg agt yca agc aat tcy ctk gcc tca gcc ccc cga ggg Leu Ser Xaa Ser Asn Ser Leu Ala Ser Ala Pro Arg Gly 20 25 30	246
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<222> 320 <223> n=a, g, c or t

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ccg aca gca cgg aag cag ctg gat aaa gaa cag gtt aga aag gca gtg Pro Thr Ala Arg Lys Gln Leu Asp Lys Glu Gln Val Arg Lys Ala Val -65 -55	150										
gac gct ctc ttg acg cat tgc aag tcc agg aaa aac aat tat ggg ttg Asp Ala Leu Leu Thr His Cys Lys Ser Arg Lys Asn Asn Tyr Gly Leu -50 -45 -40 -35	198										
ctt ttg aat gag aat gaa agt tta ttt tta atg gtg g	246										
att cca agt aaa gaa ctg agg gtc aga ttg acc ttg cct cat agt att Ile Pro Ser Lys Glu Leu Arg Val Arg Leu Thr Leu Pro His Ser Ile -15 -10 -5	294										
cga tca gat tca gaa gat atc tgt tna ttt acg aag gat gg Arg Ser Asp Ser Glu Asp Ile Cys Xaa Phe Thr Lys Asp  1 5 10	335										
<210> 659 <211> 197 <212> DNA <213> Homo sapiens <220> <221> CDS											
<222> 20196	٠.										
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acg aac aca gga aca ttg ata gag cta aat ctg mcc agc cct gta gcc Thr Asn Thr Gly Thr Leu Ile Glu Leu Asn Leu Xaa Ser Pro Val Ala -15 -10 -5	100										
ctc cag tgg cca ctt tcc agc ccc tct tgc ctg agg atc ctc agc aac Leu Gln Trp Pro Leu Ser Ser Pro Ser Cys Leu Arg Ile Leu Ser Asn 1 5 10	148										
aag gtg ccc agg aac ctg agg tgg cag aaa cac tac tcc acc cac cag g Lys Val Pro Arg Asn Leu Arg Trp Gln Lys His Tyr Ser Thr His Gln 15 20 25 30	197										
<210> 660 <211> 272											

<212> DNA

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<400> 660

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<210> 661

<211> 411

<212> DNA

<213> Homo sapiens

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<222> 263..409

<221> sig_peptide

<222> 263..340

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seq WGNLSFHLQEAHG/SE

Phe Ser Leu Glu Glu Trp Ser Leu

## <400> 661

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Ser Glu Ile Ala Glu Met Gly Ala Gly Ile Leu Glu Glu Lys Asn Tyr

1 10 15

ggy caa caa wat cac tgt aac ta

ggv caa caa wat cac tgt aac ta 411 Gly Gln Gln Xaa His Cys Asn

20

<210> 662

<211> 146

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                                       Met Ser Leu Pro Pro Phe
                                       -30
103
Phe His Pro Ser Pro Ala Pro Ser Leu Ala Pro Pro Pro Ser Leu Phe
               -20
                                  -15
ctt tcc ctc cct ccc tct ctt tct ccc cct cta ccc gcc cgg g
                                                                  146
Leu Ser Leu Pro Pro Ser Leu Ser Pro Pro Leu Pro Ala Arg
                              1
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      seq MFFLCGFLYLCFI/SF
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caatatgct atg ttt ttc ctt tgt ggt ttt ctg tat cta tgt ttt atc tca
                                                                   51
         Met Phe Phe Leu Cys Gly Phe Leu Tyr Leu Cys Phe Ile Ser
                     -10
ttt ttt ttt ttt tt
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Phe Phe Phe
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aat c Asn L	tt g eu C	gaa d Glu 1	Leu 1	ctg Leu -10	ggc Gly	tca Ser	agt Ser	tat Tyr	aat Asn -5	ccc Pro	atc Ile	tca Ser	gcc Ala	tct Ser 1	cca Pro		103
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358

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Ile Cys Phe Phe Arg Leu Thr Ile Leu Xaa Phe His Asp Asn Thr Trp

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cca ctc tat ggg ata atg cta att ttt ttc cct aaa gtg gtt tat aac Pro Leu Tyr Gly Ile Met Leu Ile Phe Phe Pro Lys Val Val Tyr Asn -40 -35 -30	L50
Asn Gln Pro Leu His Tyr Lys Ser Val Met Val Phe Gln Leu Thr Ser -25 -20 -15	198
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tgg tac atg gta gta aac cta ttg atg gca nbt ttg ctg act gtg gaa  Trp Tyr Met Val Val Asn Leu Leu Met Ala Xaa Leu Leu Thr Val Glu -20 -15 -10 -5	22
gtg act cat cca aac tcc atg cca gct gtc aac att cag tat gaa gtc Val Thr His Pro Asn Ser Met Pro Ala Val Asn Ile Gln Tyr Glu Val	70

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447

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tgc gcc tgg tgg ctg ctt ctc cca gtt tgg aag ctg gga ggg cag ctt Cys Ala Trp Trp Leu Leu Leu Pro Val Trp Lys Leu Gly Gly Gln Leu -10 -5 1	145
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58

Met Asn

367 -65 -60 -55 act aca aac caa acc aat gga tct tct act aca gga gat aaa cct gtt 154 Thr Thr Asn Gln Thr Asn Gly Ser Ser Thr Thr Gly Asp Lys Pro Val -40 gaa tca atg cag aca aaa ttg aac tac ctt aga aga aat cta ctc att 202 Glu Ser Met Gln Thr Lys Leu Asn Tyr Leu Arg Arg Asn Leu Leu Ile -25 -30 tta gtt ggt att atc atc atg gtt ttt gtc ttt atc tgt ttt tgt tat 250 Leu Val Gly Ile Ile Ile Met Val Phe Val Phe Ile Cys Phe Cys Tyr -15 -10 ctc cat tat aat tgt ctg agc gat gat gcg tcc aaa gca gga atg gtc 298 Leu His Tyr Asn Cys Leu Ser Asp Asp Ala Ser Lys Ala Gly Met Val aag aaa aaa ggc ata gca gcc aag tca tct aaa aca tca ttc agt gaa 346 Lys Lys Lys Gly Ile Ala Ala Lys Ser Ser Lys Thr Ser Phe Ser Glu gcc aag aca gcc tct caa tgc agt tca gaa aca caa acc ggg 388 Ala Lys Thr Ala Ser Gln Cys Ser Ser Glu Thr Gln Thr Gly <210> 691 <211> 408 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 304..408 <221> sig_peptide <222> 304..387 <223> Von Heijne matrix score 3.59999990463257 seq IFFSLTLSGCKFS/KL <400> 691 cttgacttct gtgcactcac aggcttgatc aacaccacaa ggaagctgcc aaggccatcc 60 tetgaaacca cageeegage tetatgttgg ceeettttag ceatggetgg aatggetgag 120 acacaggaca ccaagteeet aggetgtaca cageaetggg accetgggee etgeecatgg 180 aacaattttt teeteetaaa tetteaggee tgtgatggga ggggetaeeg caaaggtete 240 tgacatgccc cagatacatt ttccctattg tcttggggat taacatttgg ctcctcgtta 300 ctt atg caa att tct gca gcc agc ttg aat ttc tcc tca aaa aat gga 348 Met Gln Ile Ser Ala Ala Ser Leu Asn Phe Ser Ser Lys Asn Gly -20 att ttc ttt tct tta aca ttg tca ggc tgc aaa ttt tcc aaa ctt tta 396 Ile Phe Phe Ser Leu Thr Leu Ser Gly Cys Lys Phe Ser Lys Leu Leu -5 1 tgc cct ttt ggg 408 Cys Pro Phe Gly 5 <210> 692 <211> 322 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 106..321 <221> sig_peptide <222> 106..261 <223> Von Heijne matrix

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ggt cgc gac aag gag gag gag gag gtg gcc ggt gga gac tgc ata Gly Arg Asp Lys Glu Glu Glu Glu Val Ala Gly Gly Asp Cys Ile  -35 -30 -25	100
ggg agc acg gtc tac agc aaa cac tgg ctc ttc ggc gtc ctc agc gga Gly Ser Thr Val Tyr Ser Lys His Trp Leu Phe Gly Val Leu Ser Gly -20 -15 -10	148
ctc akc cag rtt gtt agc cct gga aaa cac caa aat cta ggc tca grt Leu Xaa Gln Xaa Val Ser Pro Gly Lys His Gln Asn Leu Gly Ser Xaa -5 1 5	196
gmt gag gag cag ctg acg gag ctt gat gaa cga aat gg Xaa Glu Glu Leu Thr Glu Leu Asp Glu Arg Asn 10 15 20	234
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ttg gct gga tat gaa att ctt ggt tgt cat ttc ttt tct tta gca ctg Leu Ala Gly Tyr Glu Ile Leu Gly Cys His Phe Phe Ser Leu Ala Leu -15 -10 -5	403
cta aat aca ggc ccc caa tat ctt ttg gct tat agg gtt tct gct gaa Leu Asn Thr Gly Pro Gln Tyr Leu Leu Ala Tyr Arg Val Ser Aia Glu 1 5 10	451
agg t Arg	455
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ggg gtc gcg ccg tgg cgg agc agc ctc cat ccc tgt gag atc act g Gly Val Ala Pro Trp Arg Ser Ser Leu His Pro Cys Glu Ile Thr A -25 -20 -15	Ala
ctg agc caa tcc cta cag ccc tta cgg aag ctg cct ttt aga gcc t Leu Ser Gln Ser Leu Gln Pro Leu Arg Lys Leu Pro Phe Arg Ala S -10 -5 1	
ygc acg gg Xaa Thr	153
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•	Met
gcg ccc aag ggc aaa gtg ggc acg aga ggg aag aag cag ata ttt g Ala Pro Lys Gly Lys Val Gly Thr Arg Gly Lys Lys Gln Ile Phe G -45 -40 -35	lu
gag aac aga gag act ctg aag ttc tac ctg cgg atc ata ctg ggg g Glu Asn Arg Glu Thr Leu Lys Phe Tyr Leu Arg Ile Ile Leu Gly A -30 -25 -20	Ala
aat gcc att tac tgc ctt gtg acg ttg gtc ttc ttt tac tca tct g Asn Ala Ile Tyr Cys Leu Val Thr Leu Val Phe Phe Tyr Ser Ser A -15 -5	Ala
tca ttt tgg gcc tgg ttg gcc ctg ggc ttt agt ctg gca gtg tat g Ser Phe Trp Ala Trp Leu Ala Leu Gly Phe Ser Leu Ala Val Tyr G 1 5 10 15	
gcc agc tac cac tct atg agc tcg atg gca cga gcg gcg ttc tct g Ala Ser Tyr His Ser Met Ser Ser Met Ala Arg Ala Ala Phe Ser G 20 25 30	
gat ggg gcc ctg atg gat ggt ggc atg gac ctc aac atg gag cag g Asp Gly Ala Leu Met Asp Gly Gly Met Asp Leu Asn Met Glu Gln G 35 40 45	
atg gca gag cac ctt aag gat gtk atc cta ctg aca gcc atc gtg c	ag 454

371 Met Ala Glu His Leu Lys Asp Val Ile Leu Leu Thr Ala Ile Val Gln 55 gtg ctc agc tgc ttc tct ctc tat gtc tgg tcc ttc tgg 493 Val Leu Ser Cys Phe Ser Leu Tyr Val Trp Ser Phe Trp 70 <210> 698 <211> 174 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 8..172 <221> sig_peptide <222> 8..94 <223> Von Heijne matrix score 3.59999990463257 seq AFNKAVWFTPCSC/QE <400> 698 aacaaag atg gcg gcg gtg act gtg acg gtg acg aag acg gcg gcg gcg 49 Met Ala Ala Val Thr Val Thr Val Thr Lys Thr Ala Ala Ala -25 -20 geg acg gca ttt aac aag gcg gtg tgg ttt act cca tgc agt tgt cag 97 Ala Thr Ala Phe Asn Lys Ala Val Trp Phe Thr Pro Cys Ser Cys Gln -15 -5 -10 gag gta agt agc agg ctg ccg gct cgg acg gcg gcg acg cag gac 145 Glu Val Ser Ser Arg Leu Pro Ala Arg Thr Ala Ala Thr Arg Gln Asp 10 15 agg gcg gat aag aag gag cgg ccc tgt gg 174 Arg Ala Asp Lys Lys Glu Arg Pro Cys 20 <210> 699 <211> 300 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 199..300 <221> sig_peptide <222> 199..255 <223> Von Heijne matrix score 3.59999990463257 seq PGSAICLWHSTLG/GX <221> misc feature <222> 261 <223> n=a, g, c or t <400> 699 attiliytete qqeaqeqqtq geegwagete categoattt tatgitictg yeyagaayyy 🗀 aacqqaqttt tcatcaqqta qattqqtttt trtqcqqccq tcctccaccq tttcctccaq 120 gacagcacct agtcgtggcc ggaggagtct catagctgtc agaaagaata agactgattt 180 tatgggaaaa ttaagcag atg ctc cag ttt gag aaa cct gga tct gcg atc 231 Met Leu Gln Phe Glu Lys Pro Gly Ser Ala Ile -10 -15

tgt ttg tgg cac agc act ttg gga ggy ymn ggc ggg cgt gag att gds

Cys I	Նeu	Trp	His -5	Ser	Thr	Leu	Gly	Gly 1	Xaa	Gly	Gly	Arg 5	Glu	Ile	Xaa	
agt t Ser 1	_	_	cca	_	_			•				,				300
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cta a Leu l															gg	159
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acc of												acc	cag			105
gat (	gcc					ctg					ctc-					153
tyt ( Cys )				tct					gtg					gci		201
aac (																249
tcg (	_	_		_				g								274

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                                                                       115
                                                    Met Pro Ala
tgc ctt tct tcc ttt gtc att ccc tct ctc ctt tct ccc tcc tcc cct
                                                                       163
Cys Leu Ser Ser Phe Val Ile Pro Ser Leu Leu Ser Pro Ser Ser Pro
    -10
                        -5
ccc tcc ata ggg
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Pro Ser Ile Gly
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                                                                       120
tctggatatt agccctttgt cagatgagta gattgtaaaa attttctccc attctacagg
                                                                       180
ttgcctgttc actctg atg gta gtt tct ttt gct ggt tct tgc aca att cta
                                                                       232
                  Met Val Val Ser Phe Ala Gly Ser Cys Thr Ile Leu
                      -15
ggc gcc agt agc cat tca ttc ccc att gaa gtc agc ctg ttc cca gtg
                                                                       280
Gly Ala Ser Ser His Ser Phe Pro Ile Glu Val Ser Leu Phe Pro Val
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gac tgt ggc ttc ctc ttg
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cat agc cac ctg agc ctg gtg ggg cac agc agg gcc tgt gga gtc aca His Ser His Leu Ser Leu Val Gly His Ser Arg Ala Cys Gly Val Thr -10 -5 1	343
tcc cgg cct cat gct cgg cat agg gga cgc tgc tta ggt cca tgc agt Ser Arg Pro His Ala Arg His Arg Gly Arg Cys Leu Gly Pro Cys Ser 5 10 15 20	391
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ccc tca ctt ctg aac cac cct gct tcc agc ctc atc tcc cat gat ccc Pro Ser Leu Leu Asn His Pro Ala Ser Ser Leu Ile Ser His Asp Pro -20 -10	280
tgg cca cgc ggt gcg ttt gcg ctt tca tgt cca agt gct tcc ttc atg Trp Pro Arg Gly Ala Phe Ala Leu Ser Cys Pro Ser Ala Ser Phe Met -5	328
ttg ttt tct tcc tta caa tgc cct ttc cct tat tgd naa aca gag tgc Leu Phe Ser Ser Leu Gln Cys Pro Phe Pro Tyr Xaa Xaa Thr Glu Cys 10 20 25	376
aac gwg	382

Asn xaa	
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-15 ctc ttc cgg gct gta gct gac cag gtg tat gga gac cag gac atg cat	283
Leu Phe Arg Ala Val Ala Asp Gln Val Tyr Gly Asp Gln Asp Met His -10 -5 1 5	203
gag gtt gtg cga aag cat tre atg gae tat ctg atg aag aat gee gae	331
Glu Val Val Arg Lys His Xaa Met Asp Tyr Leu Met Lys Asn Ala Asp 10 15 20	
tay tto too are tat gto aca gag gac ttt acc acc tac att akc agg	379
Tyr Phe Ser Xaa Tyr Val Thr Glu Asp Phe Thr Thr Tyr Ile Xaa Arg 25 30 35	
aag cg Lys	384
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ggctcagttt gatat atg gtt cac ctc att ctt act gaa gtc ctc att atg Met Val His Leu Ile Leu Thr Glu Val Leu Ile Met	111
atc akc gag gcn nsg aat gtg tgg tgt ggg gat tcg gg	149
Ile Xaa Glu Ala Xaa Asn Val Trp Cys Gly Asp Ser	

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377

-5

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Ser Ser His Ser Pro Thr Cys Ala Cys Lys Leu

-5

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                                                                      120
ggctgccagt agcaaaaatc actgtaattc aaaaagcatg acactacggt agtgaaatta
                                                                      180
tcacactttt ctttgcatag agcagtttac ttgtg atg att ttc aaa gat gtg
                                                                      233
                                       Met Ile Phe Lys Asp Val
                                       -20
tte tee cae ttg tea ggt tea tet ett caa etg tgt gte gea caa ttt
                                                                      281
Phe Ser His Leu Ser Gly Ser Ser Leu Gln Leu Cys Val Ala Gln Phe
                -10
                                    -5
ctc gaw ctc agt gct gtt gac at
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Leu Xaa Leu Ser Ala Val Asp
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aatactatag gtttttgctt tgtggttaac atg aag ctt aca aaa aat atc tta
                                                                      114
                                 Met Lys Leu Thr Lys Asn Ile Leu
                                                  -40
twa gta ata ata ggc tgt ttt aag ctg ata gcc tac aaa aac tct gta
                                                                      162
Xaa Val Ile Ile Gly Cys Phe Lys Leu Ile Ala Tyr Lys Asn Ser Val
                        -30
                                            -25
ctg tac ttt tac tct aac ttc tca ttt tct ttt ctt ttc ttt ttc
                                                                      210
Leu Tyr Phe Tyr Ser Asn Phe Ser Phe Ser Fhe Leu Phe Phe Phe Phe
-20
                    -15
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otto ted the ter ted tet the fee te
                                                                      242
Leu Ser Phe Phe Phe Phe Phe Phe Phe
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cag caw rag cca acg cta tca ggc cag cgt ttt aaa act aga aaa aga	162
Gln Xaa Xaa Pro Thr Leu Ser Gly Gln Arg Phe Lys Thr Arg Lys Arg -80 -75 -70	
gat gaa aaa gag agg ttt gac cct act cag ttt caa gac tgt att att Asp Glu Lys Glu Arg Phe Asp Pro Thr Gln Phe Gln Asp Cys Ile Ile -65 -55	210
caa ggc tta act gaa acc ggt act gat ttg gaa gca gta gct aag ttt Gln Gly Leu Thr Glu Thr Gly Thr Asp Leu Glu Ala Val Ala Lys Phe	258
-50 -45 -40 -35 ctt gat gct tct gga gca aaa ctt gat tac cgt cga tat gca gaa aca	306
Leu Asp Ala Ser Gly Ala Lys Leu Asp Tyr Arg Arg Tyr Ala Glu Thr -30 -25 -20	
ctc ttt gac att ctg gtg gct ggt kga atg ctg gcc cca ggt ggt aca	354
Leu Phe Asp Ile Leu Val Ala Gly Xaa Met Leu Ala Pro Gly Gly Thr -15 -10 -5	
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atagtttcaa aatcttttat ctccaggttt gatctctctt gtgaactctg gaactgtatt	180
cocaattgtc aattggacat coctacgtat gggacctcag atatttcaaa catgatgtgt	240
ccaagtetgt atcacttetg gecatestat rgttetttta ftfttteeaaa ttteacatea ccagtaacaa actagetgtg atc atg gea gat age etg gaa ata aaa ete eee Met Ala Asp Ser Leu Glu Ile Lys Leu Pro -15	307 353
ttt tta ccc ttt gca cag caa att gac atc aaa tcc tgt ttc tac ttt Phe Leu Pro Phe Ala Gln Gln Ile Asp Ile Lys Ser Cys Phe Tyr Phe -5 1 5	401
ttt ttt ttw aac wat kgc ttc cct agg g	429

Phe Phe Xaa Asn Xaa Xaa Phe Pro Arg <210> 718 <211> 350 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 4..348 <221> sig peptide <222> 4..108 <223> Von Heijne matrix score 3.5 seq ATAAATAASATTG/AS <221> misc_feature <222> 155 <223> n=a, g, c or t <400> 718 tga atg gac aga aaa tgg acc tgg aag aga ggg caa agg tca cat ctg 48 Met Asp Arg Lys Trp Thr Trp Lys Arg Gly Gln Arg Ser His Leu -30 -25 gag toa ggc cag gct gcc ccg gcc act gca gca gct acg gca gca tct 96 Glu Ser Gly Gln Ala Ala Pro Ala Thr Ala Ala Thr Ala Ala Ser -15 -10 gcc aca acg ggg gca agt gtg tgg aga agc aca atg ggc wac ctg tgt 144 Ala Thr Thr Gly Ala Ser Val Trp Arg Ser Thr Met Gly Xaa Leu Cys gat tgc acc and dca cct tat gaa ggg ccc ttt tgc aaa aaa gag gtt 192 Asp Cys Thr Xaa Xaa Pro Tyr Glu Gly Pro Phe Cys Lys Lys Glu Val 20 tet get gtt ttt gag get gge acg teg gtt act tac atg ttt caa gaa 240 Ser Ala Val Phe Glu Ala Gly Thr Ser Val Thr Tyr Met Phe Gln Glu 35 ccc tat cct gtg acc aag aat ata agc ctc tca tcc tca gct att tac 288 Pro Tyr Pro Val Thr Lys Asn Ile Ser Leu Ser Ser Ser Ala Ile Tyr 50 55 aca gat tca gct cca tcc aag gaa aac att gca ctt agc ttt gtg aca 336 Thr Asp Ser Ala Pro Ser Lys Glu Asn Ile Ala Leu Ser Phe Val Thr acc caa gca ccg gg 350 Thr Gln Ala Pro <210> 719 <211> 305 <212> DNA <213> Homo sapiens <220> <721> CDS <222> 84..305 <221> sig_peptide <222> 84..212 <223> Von Heijne matrix score 3.5

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cca tta tcc cca gag gag ttg ttg aaa agt gga ggg gtg aat cag tat Pro Leu Ser Pro Glu Glu Leu Leu Lys Ser Gly Gly Val Asn Gln Tyr -30 -25 -20	161
gtt gtg caa gag gta ctg tcc atc aaa cat ctt cca cca cag ctt aga Val Val Gln Glu Val Leu Ser Ile Lys His Leu Pro Pro Gln Leu Arg -15 -10 -5	209
gct ttt cag gct gcc ttt cga gct cag ggg ccc ctg gct atg ctg cag Ala Phe Gln Ala Ala Phe Arg Ala Gln Gly Pro Leu Ala Met Leu Gln 1 5 10 15	257
cac ttt gat act atc tac agc att ttg cat cac ttt cga agt ata gat His Phe Asp Thr Ile Tyr Ser Ile Leu His His Phe Arg Ser Ile Asp 20 25 30	305
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aaaag atg gct gct gtg caa gtt gtc ggt tcg tgg cct tcc gtg cag ccg Met Ala Ala Val Gln Val Val Gly Ser Trp Pro Ser Val Gln Pro	50 98
aaaag atg gct gct gtg caa gtt gtc ggt tcg tgg cct tcc gtg cag ccg  Met Ala Ala Val Gln Val Val Gly Ser Trp Pro Ser Val Gln Pro  -15  -10  -5  cgg gag gca ccg cgg gaa gca atc cct gag cga ggc aat ggg ttt cgc Arg Glu Ala Pro Arg Glu Ala Ile Pro Glu Arg Gly Asn Gly Phe Arg  1  5  10  15  ctc ttg tct gcc agg ctc tgc gcc ctg cgc ccg gat gac agc agc tcc Leu Leu Ser Ala Arg Leu Cys Ala Leu Arg Pro Asp Asp Ser Ser Ser  20  25  30	
aaaag atg gct gct gtg caa gtt gtc ggt tcg tgg cct tcc gtg cag ccg  Met Ala Ala Val Gln Val Val Gly Ser Trp Pro Ser Val Gln Pro  -15  -10  -5  cgg gag gca ccg cgg gaa gca atc cct gag cga ggc aat ggg ttt cgc Arg Glu Ala Pro Arg Glu Ala Ile Pro Glu Arg Gly Asn Gly Phe Arg  1  5  10  15  ctc ttg tct gcc agg ctc tgc gcc ctg cgc ccg gat gac agc agc tcc Leu Leu Ser Ala Arg Leu Cys Ala Leu Arg Pro Asp Asp Ser Ser	98
aaaag atg gct gct gtg caa gtt gtc ggt tcg tgg cct tcc gtg cag ccg  Met Ala Ala Val Gln Val Val Gly Ser Trp Pro Ser Val Gln Pro  -15  -10  -5  cgg gag gca ccg cgg gaa gca atc cct gag cga ggc aat ggg ttt cgc Arg Glu Ala Pro Arg Glu Ala Ile Pro Glu Arg Gly Asn Gly Phe Arg  1  5  10  15  ctc ttg tct gcc agg ctc tgc gcc ctg cgc ccg gat gac agc agc tcc Leu Leu Ser Ala Arg Leu Cys Ala Leu Arg Pro Asp Asp Ser Ser Ser  20  25  30  gcc cgm acc gag atc cac ctg mtc ttc gat cag ctc atc tcc gag aac Ala Arg Thr Glu Ile His Leu Xaa Phe Asp Gln Leu Ile Ser Glu Asn	98 146
aaaag atg gct gct gtg caa gtt gtc ggt tcg tgg cct tcc gtg cag ccg  Met Ala Ala Val Gln Val Val Gly Ser Trp Pro Ser Val Gln Pro  -15  -10  -5  cgg gag gca ccg cgg gaa gca atc cct gag cga ggc aat ggg ttt cgc Arg Glu Ala Pro Arg Glu Ala Ile Pro Glu Arg Gly Asn Gly Phe Arg 1  5  10  15  ctc ttg tct gcc agg ctc tgc gcc ctg cgc ccg gat gac agc agc tcc Leu Leu Ser Ala Arg Leu Cys Ala Leu Arg Pro Asp Asp Ser Ser Ser  20  25  30  gcc cgm acc gag atc cac ctg mtc ttc gat cag ctc atc tcc gag aac Ala Arg Thr Glu Ile His Leu Xaa Phe Asp Gln Leu Ile Ser Glu Asn  35  40  45  tac agc gag ggc agt ggc gtg gcc ccg gag gac gtw agt gct ctt ctt Tyr Ser Glu Gly Ser Gly Val Ala Pro Glu Asp Val Ser Ala Leu Leu 50  55  60	98 146 194
aaaag atg gct gct gtg caa gtt gtc ggt tcg tgg cct tcc gtg cag ccg  Met Ala Ala Val Gln Val Val Gly Ser Trp Pro Ser Val Gln Pro  -15  cgg gag gca ccg cgg gaa gca atc cct gag cga ggc aat ggg ttt cgc Arg Glu Ala Pro Arg Glu Ala Ile Pro Glu Arg Gly Asn Gly Phe Arg 1 5 10 15  ctc ttg tct gcc agg ctc tgc gcc ctg cgc ccg gat gac agc agc tcc Leu Leu Ser Ala Arg Leu Cys Ala Leu Arg Pro Asp Asp Ser Ser Ser  20 25 30  gcc cgm acc gag atc cac ctg mtc ttc gat cag ctc atc tcc gag aac Ala Arg Thr Glu Ile His Leu Xaa Phe Asp Gln Leu Ile Ser Glu Asn  35 40 45  tac agc gag ggc agt ggc gtg gcc ccg gag gac gtw agt gct ctt ctt Tyr Ser Glu Gly Ser Gly Val Ala Pro Glu Asp Val Ser Ala Leu Leu  50 55 60  gtc cag gct tgc ggg Val Gln Ala Cys Gly	98 146 194 242

383

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                                                                     120
atttetttga tttttatett geagttette tattgagttt tgeatgttgg etateatgtt
                                                                     180
ttaaattttc attttcata gtattctgtc ctatgg atg ttt cat ggc tgt cat
                                                                     234
                                        Met Phe His Gly Cys His
                                        -30
att tta tct ttt ctg agg ata tca act aga ggt ttt ctt ttt ctt
                                                                     282
Ile Leu Ser Phe Leu Arg Ile Ser Thr Arg Gly Phe Leu Phe Phe Leu
                -20
                                    -15
caa ttt tcc ttt cct ctg tat tat ctc ttt cgg ngg ntt ttc cct cag
                                                                     330
Gln Phe Ser Phe Pro Leu Tyr Tyr Leu Phe Arg Xaa Xaa Phe Pro Gln
tct ttc atg ttg gag gca ttt gtc aga tgt
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Ser Phe Met Leu Glu Ala Phe Val Arg Cys
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gtt atg tat aga cat tcc aaa cag cgt aat aat gtc cca tgc ctt gta
                                                                      108
    Met Tyr Arg His Ser Lys Gln Arg Asn Asn Val Pro Cys Leu Val
        -25
                            -20
                                                -15
etc tac gcc cct tgg gtc cct ccc ctc cta gct ttc tgg ggc tgg
                                                                     156
Leu Tyr Ala Pro Trp Val Pro Pro Leu Leu Ala Phe Trp Gly Trp
    -10
                        -5
tgg ctc ctg gag cag ggt ctt ttt ttt ttt tt
                                                                     191
Trp Leu Leu Glu Gln Gly Leu Phe Phe Phe
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PCT/IB99/00712

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                                                                      120
gmat atg ttt cta act ttt ttt ttc tgc aca caa gtt cat ggt cct tct
                                                                      169
     Met Phe Leu Thr Phe Phe Cys Thr Gln Val His Gly Pro Ser
ata ctt gat agc cca gct
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Ile Leu Asp Ser Pro Ala
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                                          Met Val Thr Leu Trp Ile
                                                           -10
                                                                      104
ttt caa ttt ttc ttg tgt ttg act tgt aaa gct tat aat tta aga aac
Phe Gln Phe Phe Leu Cys Leu Thr Cys Lys Ala Tyr Asn Leu Arg Asn
            -5
tgt aat gat ggg aag ggh wga gsm tca gwg gtg ctt gga ttg gaa caa
                                                                      152
Cys Asn Asp Gly Lys Gly Xaa Xaa Ser Xaa Val Leu Gly Leu Glu Gln
mnr cta cct gaa tct gct ggt atg gta caw ttt tta ggt ttg aaa cac
                                                                      200
Xaa Leu Pro Glu Ser Ala Gly Met Val Xaa Phe Leu Gly Leu Lys His
25
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agg tgg g
                                                                      207
Arg Trp
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gtt ttg tgg gct ggg ccc akg gtc ccc ctg ctg tgt gca gcc tas gga .  Val Leu Trp Ala Gly Pro Xaa Val Pro Leu Leu Cys Ala Ala Xaa Gly  -10 -5 1	104
ctt ggt gcc ctg cat ccc aga tgc tct agt caa ggc ttg agg ctt gcr Leu Gly Ala Leu His Pro Arg Cys Ser Ser Gln Gly Leu Arg Leu Ala 5 10 15	152
sct tct gaa gcc Xaa Ser Glu Ala 20	164
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388	
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ggaggtgggt ctttgcggcc gatcgcgctc ctgggggtgt tgtaacggca  gag gtt cca Glu Val Pro 15 gag gaa gaa	atccace gggtcte gagctge ggaaag a aga ce Arg Pi	gact the cape of t	tegeace teetgag gttegtg at ggg sn Gly ecc ca Pro H:	ccc gt ctt cc gtg cc ggt ca agg g Arg A s ac ata is Ile ca gaa	tgttg gccat ggtga cggat ct ga la As ggg Gly	Jecc ttt ictg geg it ti sp Ph cct Pro	ccts cggs tgts tt cone An gat Asp	Jegta Jegta Jegga Jegga Jegga Tyr 25 gea	atc of gradients of the	cccto agggt cggag ctgct cg aa ro As cca Pro	caccac tggaca gtgctt tcagtc at gca sn Ala aca Thr	180 240 300 360 412
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Val Asp Phe Lys Ile Arg Thr Ile Glu Leu Asp Gly Lys Thr Ile Lys
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Ser Ser Tyr Tyr Arg Gly Ala His Gly Ile Ile Val Val Tyr Asp Val
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399													
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gag aaa aag tgt gct gtg gtt cgg aag tct aag cag ggc agg aaa cgc
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gcc caa ggg gtc cgg tgc tca gac tgt gga ttg aac gta
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Pro Ser Xaa Ser Ala Val Va -15 -1	-5	
aat cca tca tct ccc tat ac Asn Pro Ser Ser Pro Tyr Th 1 5	r Asn Ser Ser Arg Lys Glr 10	Pro Met Ser 15
gca aca ctt aga gaa aga tt Ala Thr Leu Arg Glu Arg Le 20	a Arg Lys Thr Arg Phe Ser 25	Phe Asn Ser
tct nac aat gtg gtg aac gt Ser Xaa Asn Val Val Asn Va 35		aatgat 249
cagacetttt cagagaacec agea agtttaaame atatagreag tgat ateteaatgt etgtgaatet eagt	ttgaag aaaatacaaa tttgnaa	aat actttgaaga 369
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-10

-15

-5

WO 99/53051 PCT/I	D00/00713
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tac ctt ctg tcg scc ctg ctc tct gmt gcc ttc cta ctc gtg agg maa Tyr Leu Leu Ser Xaa Leu Leu Ser Xaa Ala Phe Leu Leu Val Arg Xaa 15 20 25	193
ctg ccg ctc tgc cac ggt ctg ccc acc caa cgc gaa smc ggt aac Leu Pro Pro Leu Cys His Gly Leu Pro Thr Gln Arg Glu Xaa Gly Asn 30 35 40	241
ccg tcr wsa ytt tgactgggtg agcctcccgc gtgttagtac cccgcgacsk Pro Ser Xaa Xaa	293
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geteatetge aaagagataa ngaggteeet geggatgtg atg gee cag eta tgg	234
Met Ala Gln Leu Trp	
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Leu Ser Cys Phe Leu Leu Pro Ala Leu Val Val Ser Val Ala Ala Asn	
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Val Ala Pro Xaa Phe Leu Ala Asn Met Thr Ser Val Ile Leu Pro Glu	
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WO 99/53051 PCT/I	B99/00712
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aga gct ccg gca aaa cct cca ggt agt gga ttg gac ttg gct gat gct Arg Ala Pro Ala Lys Pro Pro Gly Ser Gly Leu Asp Leu Ala Asp Ala 45 50 55	304
ttg gat gat caa gat gat ggc cgc aga aac cgg gta taggaggaag Leu Asp Asp Gln Asp Asp Gly Arg Arg Asn Arg Val  60 65	350
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gag gac gac ttc aac tat ggc agc gtg gcc tcc gcc acc gtg cac Glu Asp Asp Phe Asn Tyr Gly Ser Ser Val Ala Ser Ala Thr Val His -40 -35 -30 -25	159
atc cga atg gcc ttt ctg aga aaa gtc tac agc att ctt tct ctg cag Ile Arg Met Ala Phe Leu Arg Lys Val Tyr Ser Ile Leu Ser Leu Gln	207

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-50

-45

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								gaa Glu								212
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atc					ggc			ctg Leu		gtg					aat	308
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		gct					tcc	ttc Phe				cag				404
	gga	_		_		cag		gca Ala						_	_	452
ccc						gcg Ala		sca Xaa	tgag	gcact	tg d	etgea	agcg	g		499
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								tta Leu								101
								cgg Arg								149
								ctt Leu						-	_	197
	_	_	_				_	ctt Leu 10			_		_		_	245
								2ct 2co								293
								aaa Lys				_	_			341
								gaa Glu								389

WO 99/53051 PCT/IB99/00712 416 age cac att ttg cag tee tgg atg get ttg tat ttg cae taaatcagga Ser His Ile Leu Gln Ser Trp Met Ala Leu Tyr Leu His 70 aggaaaattt ttgtaca 455 <210> 783 <211> 453 <212> DNA <213> Homo sapiens <220> <221> CDS <222> 85..168 <221> sig_peptide <222> 85..144 <223> Von Heijne matrix score 5 seq ALLSVCSTDVTTA/HA <221> misc_feature <222> 284 <223> n=a, g, c or t <400> 783 ccccttgtgg ccaagcctgg aacatcacat ctgtacgttg caatctgtgg atcagctacg 60 agactgagag aaaggaatga aagg atg gaa gaa tta caa gat cag gca ctg 111 Met Glu Glu Leu Gln Asp Gln Ala Leu -20 -15 ctg tct gtc tgt tcc acg gat gta acc aca gca cac gcg tgg ctc acg 159 Leu Ser Val Cys Ser Thr Asp Val Thr Thr Ala His Ala Trp Leu Thr -10 -5 gta cta gtg tgataaatgc ttgttacatg aaggcgtgaa cagggatgag 208 Val Leu Val aagagacttc ctggagaaac aaaaggacta acaatcagga aggggaggtg atcggggcag 268 gagtaaagtg gacachtcag ctggtcccct gggtcgtcca cccgatgtcc cccattctcc 328 ccaettggcc tececcaeag geteteggca aaggacegtg ggaggcaeet gtgacaetge 388 cetttteetg tgeagetgtt tktettette attetttea etectegtta etetttttt 448 tttca 453

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gagggagca atg gtg ggg cga gtg agg gtc tgc cgt aaa tat ccc ccg acc	291
Met Val Gly Arg Val Arg Val Cys Arg Lys Tyr Pro Pro Thr	
-45 -40 -35	220
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Trp Asn Ile Cys Cys Ala Ala Ala Ala Ala Ala Ala Ala Gly Ser Arg	
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cgg ctg ggg acc cct cag caa atc gcc att gct cgc gag ggt gac ctc Arg Leu Gly Thr Pro Gln Gln Ile Ala Ile Ala Arg Glu Gly Asp Leu -30 -25 -20	154
ctg acc aag gag cgg ctg tgc tgt ggc ctg tcc atg ttc gag gtc atc Leu Thr Lys Glu Arg Leu Cys Cys Gly Leu Ser Met Phe Glu Val Ile -15 -10 -5	202
ctg acc cgc att cgg agc tac ctg cag gac ccc atc tgg cgg ggc cca Leu Thr Arg Ile Arg Ser Tyr Leu Gln Asp Pro Ile Trp Arg Gly Pro 1 5 10	250
ccg ccc acc aat ggc gtc atg cac gtc gat gag tgt gtg gag ttc cac Pro Pro Thr Asn Gly Val Met His Val Asp Glu Cys Val Glu Phe His 15 20 25 30	298
cgg ctg tgg agc gcc atg cag ttc gtg tac tgc atc cct gtg gga acc Arg Leu Trp Ser Ala Met Gln Phe Val Tyr Cys Ile Pro Val Gly Thr 35 40 45	346
aac gag ttc aca gct gag cag tgt ttc ggc gat ggc ttg aac tgg gct Asn Glu Phe Thr Ala Glu Gln Cys Phe Gly Asp Gly Leu Asn Trp Ala 50 55 60	394
ggt tek eer kea ttg tee tge tsg gee age gee get ttg ace tgt Gly Ser Pro Xaa Leu Ser Cys Xaa Ala Ser Ser Val Ala Leu Thr Cys 65 70 75	442
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tct atc ctg tgt aac tac aag gcc atc gaa atg ccc tca cac cag acc Ser Ile Leu Cys Asn Tyr Lys Ala Ile Glu Met. Pro Ser His Gln Thr -50 -45 -40	153
tac gga ggg agc tgg aaa ttc ctg acg ttc att gat ctg gtt atc cag  Tyr Gly Gly Ser Trp Lys Phe Leu Thr Phe Ile Asp Leu Val Ile Gln  -35 -30 -25	201
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cgg gct cca tcc ctc Arg Ala Pro Ser Leu -50				
tcc tcc ccc tgg ggt Ser Ser Pro Trp Gly -30			_	Сув
cac att tct ttt cca His Ile Ser Phe Pro -15				
atc ccc cga cct cac Ile Pro Arg Pro His 1	Leu Pro Pro 5	Thr Ala Ala	Cys 10	296
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gca gtg ttt tat acc ctc ggc aat ctt gct gcg tta sca gta cat gct

Ala Val Phe Tyr Thr Leu Gly Asn Leu Ala Ala Leu Xaa Val His Ala

WO 99/53051 PCT/IB99/00712 421 20 25 tnw taatgggacc tqtqaaqcaa ctqaaqaaaa tgtttgaagc aacaagattg 410 Xaa 30 cttgcaacaa ttgttatgct tttgtgtttc gtatttaccc tgtgtgctgc tctttggtgg 470 cataagaagg gactggctgt gttattctgc atattgcagt tcttgtcaat gacctggtat 530 agectgtert acateceata tgeaagggat geagttatta aatgetgtte tteteteeta 590 aqttqaaaat cagaaacttg tggaaaagag cacttgaatg ttggtactct atqtttqqtq 650 aagtttgctt ttccccataa aacactccag gaacaactga cgtgacagtt gaagaccgtt 710 ttgtactaag tctcattttg tatactggta aaaactacat gcttgattaa accattaaat 770 gcttgtaact ttaaattcat tatgtgtcat taatatactt ttccaaagat aagattttta 830 atcact 836 <210> 791

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-45 -40 -35 ate eeg atg ate gge tta ate tge etg gge atg gge age get geg ett 242 Ile Pro Met Ile Gly Leu Ile Cys Leu Gly Met Gly Ser Ala Ala Leu -30 -25

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-10 -5 aaa gaa caa ccc gga gcc ctg gaa ccg cct gag ccc caa tgaccaatac 339 Lys Glu Gln Pro Gly Ala Leu Glu Pro Pro Glu Pro Gln

10 aagttoottg cagtttocac tgactataag aagctgaaga aggaccygdc agacttot&a 399 gccaggctgg gctgccagtg ccatgcaagc cacagccagc cagcccatcc acttcttcca 459

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PCT/IB99/00712

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Tyr Trp Leu Asp Leu Trp Leu Phe Ile Leu Phe Asp Val Val Phe

367

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Leu Phe Val Tyr Phe Leu Pro

424

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Cys Val Trp Val Cys Val Tyr Thr Val Glu Ser Lys Leu Glu Asn Ser 10 15 20	
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Phe Val Thr Phe Leu Phe Pro Pro 40 45	
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gcc ctc cat ggg ggg atg aag aca ctg ctg cca tgg aca gcc cgt gcc	283
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J JJ - We day oddood grage	Met Asn Arg Thr	gca atg 113 Ala Met
aga gcc agt cag aag gac ttt Arg Ala Ser Gln Lys Asp Phe 10	or Asn Ser Xaa Asn Gln Va	g aaa ctc 161 l Lys Leu
ttg aaa aag gat cca gga aac Leu Lys Lys Asp Pro Gly Asn 25	gar tom ago tagaactota ogo	gctatat 211
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1 net	Ala Ser Ser Gly Ala Gly A	sp Pro Leu
gat tot aag ogt gga gag goo o Asp Ser Lys Arg Gly Glu Ala F	erg ttc gct cag cgt atc gac	ccg act 221
±3	20	
cgg gag aag ctg aca ccc gag c	aa ctg cat tcc atg cgg cag	gcg gag 269
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ttg ccc agt ggc aga agg tcc t Leu Pro Ser Gly Arg Arg Ser m	20 020 000	aca tcg 317
45 45 Ng Aig Sei 1	o cr	Thr Ser
tgaccggcct aggcatcggg gccctgg	tat tagatatata	tctactcga 377
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428

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Ser Ala Met Gly Phe Thr Ala Ala Gly Ile Ala Ser Ser Ser Ile Ala

W G 77/33031	429	FC1/1B99/00/1
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	75 : act ctg cag tca ctg gga gca act g : Thr Leu Gln Ser Leu Gly Ala Thr G	
85 tcc qqa ttq acc aaq tko	90 9 atc ctg ggc tcc att ggg tct gcc a	5 .tt gcg 457
	Ile Leu Gly Ser Ile Gly Ser Ala I 105 110	
gct gtc att gcg agg ttc Ala Val Ile Ala Arg Phe 115	tac tagetecetg eccetegeee tgeagage Tyr	aag 508
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Leu Trp Asn Ile Leu Lys Leu Pro Val Gln Thr Leu Leu Gln Gly
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taaacatgat gctttgaaga catatgcatc attggctaca cttccatttt tgtctactgt
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tgttactgac aagctttttg taattgatgc tttgtattca gataatataa gcaaggaaaa
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Val Lys Arg Pro Arg Gly Val Ser Leu Thr Asn His His Phe Tyr Asp
                            10
Glu Ser Lys Pro Phe Thr Cys Leu Asp Gly Ser Ala Thr Ile Pro Phe
                        25
                                            30
Asp Gln Val Asn Asp Asp Tyr Cys Asp Cys Lys Asp Gly Ser Asp Glu
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Pro Gly Thr Ala Ala Cys Pro Asn Gly Ser Phe His Cys Thr Asn Thr
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Pro Ala Ser Val Gly Asp Arg Val Thr Ile Thr Cys Arg Ala Ser Gln
                                   20
Ser Ile Gly Ser Tyr Leu Asn Trp Tyr Gln His Lys Pro Gly His Ala
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Pro Arg Leu Leu Ile Tyr Ala Ala Thr Thr Leu Ser Arg Gly Gly Pro
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Ala Arg Phe Ser
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                       -10
                                           -5
Ala Gly Arg Arg Ser Pro Xaa Thr Pro Asp Glu Ser Thr Pro Pro Pro
                                   10
Arg Lys Lys Lys Asp Ile Arg Asp Tyr Asn Asp Ala Asp Met Ala
           20
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Arg Leu Leu Glu Gln Gly Glu Gly
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Val Gln Cys Asp Val Glu Leu Val Glu Ser Gly Gly Leu Val Gln
Pro Gly Gly Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Asn Phe
                       20
Ser Thr Tyr Glu Met His Trp Ile Arg Gln Ala Pro Cly Lys Gly Pro
                   35
                                       40
Glu Trp Val Xaa Tyr Val Ser Gly Gly Gly Gly Thr Xaa Xaa Asn Ala
               50
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Xaa Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Asn Ser
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Phe Val Tyr Leu Gln Met Asp Ser Leu Arg Val Glu Asp Thr Ala Leu

90

85

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 Pro Gly Thr Ser Leu Thr Leu Ser Cys Ala Gly Ser Gly Phe Ser Phe
     15
                         20
 Ser Asp Tyr Gly Ile His Trp Val Arg Gln Ala Pro Gly Lys Gly Leu
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                                          40
 Glu Trp Val Ala Val Ile Ser His Asp Gly Asn Asn Lys Tyr Tyr Gly
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                                      55
 Gly Ser Met Lys Gly Arg Val Thr Ile Ser Arg Asp Asn Ser Arg His
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                                      -10
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                                                  10
 Pro Ser Gly Thr Leu Ser Leu Thr Cys Thr Val Xaa Gly Xaa Xaa Ile
                         20
 Thr Asn Tyr Tyr Trp Ser Xaa Ile Arg Gln Ser Pro Gly Lys Gly Leu
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436

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Val Glu Leu Ser Ile His Trp Val Arg Gln Ser Pro Gly Lys Gly Leu

437 Glu Trp Met Gly Gly Phe Asp Leu Glu Ser Gly Glu Thr Ile Tyr Ala 55 Gln Arg Phe Gln Gly Arg Ile Thr Met Thr Glu Asp Ser Ser Ser Asp 70 Thr Ala Phe Met Glu Leu Ile Ser Leu Arg Pro Glu Asp Ala Ala Val 85 Tyr Tyr Cys Ala Thr Ile Arg Leu Pro Val Val Leu Phe Phe Ala Ala 100 Ser Gly Ala Arg Glu Pro Trp Ser Pro Ser Pro Gln Xaa Pro Arg <210> 825 <211> 37 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -18..-1 <400> 825 Met Trp Leu Pro Leu Val Leu Leu Ala Val Leu Leu Leu Ala Val -10 Leu Cys Lys Val Tyr Leu Gly Leu Phe Ser Gly Ser Ser Pro Asn Pro 5 1 Phe Ser Glu Glu Arg <210> 826 <211> 51 <212> PRT < <213> Homo sapiens <220> <221> SIGNAL <222> -25..-1 <400> 826 Met Glu Leu Ala Leu Arg Arg Ser Pro Val Pro Arg Trp Leu Leu -15 -20 Leu Pro Leu Leu Gly Leu Asn Ala Gly Ala Val Ile Asp Trp Pro -5 Thr Glu Glu Gly Lys Glu Val Trp Asp Tyr Val Thr Val Arg Lys Asp 10 Ala Tyr Met 25 <210> 827 <211> 131 <212> PRT <213> Homo sapiens <220> <221> SIGNAL

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438 Thr Asn Gly His Phe Pro Tyr Trp Phe Gln Gln Lys Pro Gly Gln Ala 35 40 Pro Arg Thr Leu Ile Ser Asp Thr Phe Asn Arg Gln Ser Ser Thr Pro 55 Ala Arg Phe Ser Gly Ser Leu Leu Gly Gly Lys Ala Val Leu Thr Leu 70 Ser Asp Ala Gln Pro Asp Asp Glu Ala Glu Tyr Tyr Cys Val Leu Ser 85 90 Tyr Ser Gly Gly Arg Pro Val Phe Gly Gly Gly Thr Lys Leu Thr Val 100 105 Leu Ser Gln 110 <210> 828 <211> 25 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -21..-1 <400> 828 Met Gln Ala Cys Met Val Pro Gly Leu Ala Leu Cys Leu Leu Gly -15 Pro Leu Ala Gly Ala Lys Pro Val Gln <210> 829 <211> 79 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -23..-1 <400> 829 Met Pro Ser Tyr Lys Val Cys Gly Val Phe Cys Leu Phe Val Cys Leu -20 -15 -10 Phe Leu Ser Gln Ser Phe Ala Phe Val Leu Gln Ala Gly Val Gln Trp Arg Asp Leu Cys Ser Leu Gln Pro Gln Leu Pro Arg Phe Gly Pro Ser 10 15 20 Ser Cys Leu Ser Leu Pro Ser Gly Trp Asp Cys Arg Arg Pro Pro Pro 30 35 Arg Leu Ala Asn Ser Cys Val Phe Gly Gly Asp Gly Val Ser Pro <210> 830 <211> 59 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -21..-1 <400> 830 Met Gly Thr Gln Glu Gly Trp Xaa Leu Leu Cys Leu Ala Leu Ser -15 -20 -10 Gly Ala Ala Glu Thr Lys Pro His Pro Ala Glu Gly Gln Trp Arg Ala 1

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Pro Val Thr Glu Ala Ala Ile Phe Tyr Glu Thr Gln Xaa Ser Leu Trp

1 10 15

Ala Glu Ser Glu His Xaa Leu Lys Thr Leu Gly Gln Cys Asp Ala Asp 20 25 30

Val Pro Gly Pro Pro Gly Asp Ser Arg Leu Pro Ala Val Gln Glu Trp
35 40 45

Gly Ala Gln Glu Pro Val His Leu Asp Ser Pro Ala Ile Lys His Gln
50 55 60

Phe Leu Leu Thr Gly Asp Thr Gln Gly Arg Tyr Arg Cys Arg Ser Gly 65 70 75 80

Leu Ser Thr Gly Trp Xaa Gln Leu Ser Lys Leu Leu Glu Leu Thr Gly
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Pro Lys Val Leu Ala Cys Ser Leu Ala Leu Asp Gly Ala Ser

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<222> -19..-1

<400> 832

Met Leu Pro Ser Gln Leu Ile Gly Phe Leu Leu Trp Val Pro Ala
-15
-10
-5

Ser Arg Gly Glu Ile Val Leu Thr Gln Ser Pro Asp Phe Leu Ser Val

Thr Pro Lys Glu Lys Val Thr Ile Thr Cys Arg Ala Ser Xaa Ser Ile
15 20 25

Gly Ser Ser Leu Tyr Trp Tyr Gln Gln Lys Pro His Gln Ser Pro Lys
30 40 45

Leu Val Ile Lys Tyr Ala Ser Gln Ser Phe Ser Gly Val Ser Ser Arg
50 55 60

Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asn Ser
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Leu Glu Pro Gly 80

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 Trp
 Phe
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 Leu
 Leu
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 Val
 Arg
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 Gln
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 Leu
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 Glu

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 5
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 Ser
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 Pro
 Gly
 Leu
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 Lys
 Pro
 Ser
 Gly
 Thr
 Leu
 Ser
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 Ile
 Cys

10 15 20 Gly Val Ser Gly Asp Ser Val Thr Ile Ser Gly Trp Trp Ser Trp Val 35 30 Arg Gln Pro Pro Gly Lys Gly Leu Glu Trp Ile Ser Glu Ile Asp His 45 Gly Gly Asn Thr Asn Tyr Asn Pro Ser Leu Lys Ser Arg Val Xaa Ile 60 65 Ser Leu Asp Lys Ser Lys Asn Lys Phe Ser Leu Arg Leu Thr Ser Val 75 80 Thr Ala Ala Asp Thr Ala Met Tyr Xaa Cys Ala Arg Gly Gly Ala Xaa 95 Ser Ser Ser Ala Phe Asp Val Trp Gly Leu Xaa Thr Met Val Ile Ile Ser Ser Ala 120 <210> 836 <211> 139 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -19..-1 <400> 836 Met Asp Ile Leu Cys Ser Thr Leu Leu Leu Leu Thr Val Pro Ser Trp -10 -15 Val Leu Ser Gln Val Thr Leu Xaa Glu Ser Gly Pro Ala Leu Val Lys Ala Thr Gln Thr Leu Arg Leu Thr Cys Thr Phe Ser Gly Phe Ser Leu 20 Ser Thr Asn Arg Met Arg Val Ser Trp Ile Arg Gln Pro Pro Gly Lys 40 35 Ala Leu Glu Trp Leu Ala Arg Ile Asp Trp Asp Asp Tyr Lys Arg Tyr 55 Ser Thr Ser Leu Lys Thr Arg Val Thr Ile Ser Lys Asp Thr Ser Lys 70 Asn Gln Val Ile Leu Thr Met Thr Asn Val Asp Pro Ala Asp Thr Ala 85 Thr Tyr Tyr Cys Ala Arg Leu Ser Thr Ala Ala Thr Pro Gln Phe Phe 100 Asp Phe Trp Gly Gln Gly Val Leu Val Ser Val 115 <210> 837 <211> 139 <212> PRT <213> Homo sapiens . <220> <221> SIGNAL <222> -19..-1 <400> 837 Met Xaa Mis Leu Trp Phe Phe Leu Leu Val Ala Ala Pro Arg Trp -15 -10 -5 Val Leu Ser Gln Val Gln Leu Gln Glu Ser Gly Pro Gly Leu Val Lys Pro Ser Xaa Thr Leu Ser Leu Thr Cys Thr Val Ser Gly Asp Ser Ile 20 25 Ser Ser Tyr Tyr Trp Ser Trp Ile Arg Gln Pro Pro Gly Lys Gly Leu 45 35

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Glu Trp Ile Gly Tyr Ile Tyr Tyr Ser Gly Ser Thr Asn Tyr Asn Pro 50 55 Ser Leu Lys Ser Arg Val Thr Ile Ser Val Asp Thr Ser Lys Asn Gln Phe Ser Leu Lys Leu Ser Ser Val Thr Ala Ala Asp Thr Ala Val Tyr 85 Tyr Cys Ala Arg Xaa Leu Xaa Tyr Tyr Asp Arg Ser Gly Tyr Phe Arg 100 Tyr Phe Asp Tyr Trp Gly Gln Gly Thr Trp Ser <210> 838 <211> 136 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -19..-1 <400> 838 Met Lys His Leu Trp Phe Phe Leu Leu Leu Val Ala Ala Pro Arg Trp -15 -10 Val Leu Ser Gln Val Gln Leu Gln Glu Ser Gly Pro Gly Leu Val Lys Pro Ser Gln Thr Leu Ser Leu Thr Cys Thr Val Ser Gly Gly Ser Ile 20 Asp Ser Gly Asn Tyr Tyr Trp Ser Trp Ile Arg Gln Pro Ala Gly Lys 35 40 Gly Leu Glu Trp Ile Gly Arg Ile Tyr Ser Thr Gly Ser Thr Asn Tyr 55 Asn Pro Ser Leu Ser Ser Arg Val Gln Ile Ser Leu Asp Thr Ser Lys Asn Leu Leu Ser Leu Asn Leu Thr Ser Val Thr Ala Ala Asp Thr Ala 85 Val Tyr Phe Cys Ala Arg Thr Phe Pro Phe Tyr Trp Tyr Leu Asp Leu 100 Trp Gly Arg Gly Ile Leu Val Thr <210> 839 <211> 143 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -19..-1 <400> 839 Met Lys His Leu Trp Phe Phe Leu Leu Val Ala Ala Pro Arg Trp -10 Val Leu Ser Gln Val Gln Leu Gln Glu Ser Gly Pro Arg Leu Val Lys Pro Ser Gln Thr Leu Ser Leu Thr Cys Thr Val Ser Gly Gly Ser Ile 20 . .. 25 Ser Ser Gly Gly Tyr Phe Trp Ser Trp Ile Arg Gln His Pro Gly Arg 35 40 Gly Leu Glu Trp Ile Gly Tyr Ile Tyr Tyr Asn Trp Ser Thr Tyr Tyr 55 Asn Pro Ser Leu Arg Ser Arg Val Thr Met Ser Met Asp Thr Ser Lys

65 70 75
Asn Gln Phe Ser Leu Asn Leu Asn Ser Val Thr Ala Ala Asp Thr Xaa

443 80 85 90 Met Tyr Tyr Cys Ala Arg Gly Arg Gly Arg Leu Gly Trp Phe Xaa Xaa 100 105 Xaa Gly Xaa Gly Xaa Pro Gly His Arg Leu Ile Ser Arg Pro Gly <210> 840 <211> 111 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -19..-1 <400> 840 Met Lys His Leu Trp Phe Phe Leu Leu Val Ala Ala Pro Arg Trp -15 -10 Val Leu Ser Gln Val Gln Leu Gln Glu Ser Gly Pro Gly Leu Val Lys Pro Ser Glu Thr Leu Ser Leu Thr Cys Thr Val Ser Gly Gly Ser Ile 20 Arg Thr Gly Ser Tyr Tyr Trp Thr Trp Val Arg Gln Pro Pro Gly Lys Gly Leu Glu Trp Ile Gly Tyr Ile Tyr Tyr Thr Gly Asp Thr Tyr Tyr 55 Asn Pro Ser Leu Lys Ser Arg Ile Thr Met Ser Leu Asp Thr Xaa Xaa 70 Asn Gln Phe Xaa Leu Ser Leu Thr Ser Val Thr Val Ala Asp Thr <210> 841 <211> 53 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -15..-1 Met Lys Leu Ser Val Cys Leu Leu Val Thr Leu Ala Leu Cys Cys -10 -5 Tyr Gln Ala Asn Ala Glu Phe Cys Pro Ala Leu Val Ser Glu Leu Leu 10 Asp Phe Phe Phe Ile Ser Glu Pro Leu Phe Lys Leu Ser Leu Ala Lys 20 25 Phe Asp Ala Pro Arg 35 <210> 842 <211> 23 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -16..-1 <400> 842 Met Ser Pro Val Leu Leu Val Leu Ser Leu Ser Gln Cys Leu Leu Ser -10 Asp Pro Val Ile Pro Gly Leu

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                                   -10
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Pro Ser Glu Xaa Leu Ser Leu Thr Cys Ser Val Ser Gly Val Ser Val
                        20
Thr Asn Phe Phe Trp Asn Trp Ile Arg Lys Pro Pro Gly Lys Gly Leu
                   35
                                        40
Glu Trp Leu Gly Tyr Met Ser Tyr Gly Val Ser Thr Asn Tyr His Pro
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                                   55
Ala Tyr Gln Ser Arg Val Ser Ile Ser Ile Asp Thr Trp
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                                   -10
Val Leu Ser Gln Val Gln Leu Gln Glu Ala Gly Pro Arg Leu Val Lys
                     . 5
Pro Ser Glu Ala Leu Ser Leu Thr Cys Thr Val Ser Gly Val Ser Ser
                       20
Ser Asn Tyr Asp Trp Ser Trp Ile Arg Gln Ala Pro Gly Lys Gly Leu
Glu Trp Ile Gly Tyr Ile Asp Asp Ser Lys Asn Arg Gly Ser Thr Thr
                                    55
Tyr Asn Pro Ser Leu Lys Ser Arg Val Thr Ile Ser Xaa Asp Thr Ser
                                70
Lys Xaa Gln Leu Ser Leu Arg Leu Thr Ser Val Thr Xaa Ala Asp Thr
                            85
Ala Val Tyr Tyr Cys Ala Arg Lys Ser Ser Met His Ser Ser Gly Trp
                       100
His Asn Arg Ser Leu Tyr Trp Tyr Phe Asp Pro
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446 Pro Gly Ala Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Xaa Phe 20 Thr Xaa Xaa Ala Xaa His Trp Val Arg Gln Ala Pro Gly Gln Arg Leu 35 Glu Trp Met Gly Trp Ile Asn Ala Ala Xaa Gly Xaa Thr Xaa Tyr Ser 55 Gln Xaa Phe Gln Xaa Arg Val Thr Xaa Thr Arg Asp Thr Ser Ala Ser 70 Thr Val Ser Met Glu Leu Ser Ser Leu Arg Ser Glu Asp Thr Ala Val 85 Tyr Phe Cys Ala Arg Asp Trp Glu Ile Ala Val Val Pro Thr Ala Ile 100 Asn Ser Tyr Gly Phe Asp Pro Gly Ala Arg Glu Pro 115 <210> 848 <211> 52 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -26..-1 <400> 848 Met Glu Ala Arg Val Glu Arg Ala Val Gln Lys Arg Gln Val Leu Phe -20 Leu Cys Val Phe Leu Gly Met Ser Trp Ala Gly Ala Glu Pro Leu Arg -5 Tyr Phe Val Ala Glu Glu Thr Glu Arg Gly Thr Xaa Leu Thr Asn Leu 10 Ala Lys Asp Leu 25 <210> 849 <211> 134 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -19..-1 <400> 849 Met Asp Trp Thr Trp Ser Ile Leu Phe Leu Val Ala Ala Ala Thr Gly -15 -10 Ala His Ser Gln Val Gln Leu Val Gln Ser Gly Glu Val Lys Lys Pro Gly Ala Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe 20 Thr Arg Tyr Asp Ile Asn Trp Val Arg Gln Ala Pro Gly Gln Gly Leu 35 40 Glu Trp Met Gly Trp Ile Ser Ala Xaa Asn Gly Asn Thr Asn Tyr Ala 55 Gln Xaa Val Gln Gly Arg Val Thr Met Thr Tnr Asp Thr Ser Thr Arg 70 Thr Ala Tyr Met Glu Leu Arg Ser Leu Arg Ser Asp Asp Thr Ala Ile 85 90 Tyr Tyr Cys Ala Arg Glu Ile Xaa Val Xaa Xaa Cys Asp Gly Gln Leu Gly Pro Gly Asn Leu Val 115

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-10 -15 Ala Leu Ser Gln Val Gln Leu Val Gln Ser Gly Glu Val Lys Lys Pro Gly Ala Ser Val Arg Val Ser Cys Lys Ala Ser Gly Tyr Ser Phe Ile Gly Tyr Tyr Val His Trp Ile Arg Gln Thr Pro Gly Arg Xaa Leu 448

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Ile Cys Lys Gly Phe Leu Pro Val Tyr Leu Leu Val Leu Ser Leu
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Ser Leu Ser Leu Cys Cys Ser Leu Leu Ser Leu
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Val His Ser Gln Val His Leu Val Gln Ser Gly Ala Glu Val Lys Lys
Pro Gly Thr Pro Val Asn Ile Ser Cys Lys Ala Phe Gly Tyr Thr Phe
                       20
Pro Ala Phe Ala Ile His Trp Val Arg Gln Ala Pro Gly Gln Ser Leu
                   35
                                        40
Glu Trp Met Gly Trp Val Asn Ile Gly His Gly Asn Thr Lys Tyr Ser
                50
                                    55
Gln Lys Phe Gln Gly Arg Leu Ala Ile Ser Arg Asp Thr Ser Ala Asn
                               70
Ile Val Tyr Xaa Glu Leu Ser Gly Leu Arg Ser Glu Asp Thr Ala Val
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                                                90
Tyr Tyr Cys Ala Arg Asp Asn Leu Phe Phe Gly Ser Met Gly Phe Asp
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Met Ala Trp Thr Val Leu Leu Gly Leu Leu Ser His Cys Thr Gly
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Ser Val Thr Ser Tyr Val Leu Thr Gln Pro Pro Ser Val Ser Val Ala
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Pro Gly Lys Thr Ala Ser Ile Thr Cys Gly Gly Asp Asn Ile Glu Ser

449 20 Gln Val Val His Trp His Gln Gln Lys Pro Gly Gln Ala Pro Ile Leu 40 Val Ile Tyr Asp Asp Thr Asp Arg Pro Ser Gly Ile Pro Asp Arg Phe 55 Ser Gly Ser Asn Ser Gly His Thr Ala Thr Leu Thr Ile Ser Arg Val 75 Glu Ala Gly Asp Glu Ala Asp Tyr Tyr Cys Gln Val Trp Asp Arg Ser 90 Ser Gly Gln Gly Ile Phe Gly Gly Gly Thr Lys Leu Thr Val Leu Arg 100 105 Gln Pro Lys Ala Ala Pro Ser Val Thr Leu Phe Pro Pro Ser Ser Glu 115 120 Glu Leu Gln Ala Asn Lys Ala Thr 130 <210> 856 <211> 48 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -15..-1 <400> 856 Met Arg Leu Leu Phe Leu Leu Phe Val Cys Phe Ser Arg Gln Gly -10 -5 Leu Ala Leu Ser Leu Arg Leu Glu Cys Ser Gly Met Ile Met Ala Tyr 10 Cys Ser Ile Ser Leu Pro Gly Ser Ser Pro Leu Thr Ser Ala Ser <210> 857 <211> 74 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -19..-1 <400> 857 Met Lys His Leu Trp Phe Phe Leu Leu Leu Val Ser Ala Pro Arg Trp -15 -10 Val Leu Ser Gln Val Gln Leu Gln Glu Ser Gly Pro Gly Leu Val Lys Pro Ser Gly Arg Leu Ser Leu Ala Cys Asp Val Val Glu Leu Ser Pro 20 25 Pro Ala Pro Arg Gly Gly Ser Ala Val His Leu Arg Asn Leu Ser Ser 35 Trp Glu Pro His Leu Gln Pro Val Ser Gly 50 <210> 858 <211> 57 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -32..-1

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                            -25
                                                -20
Gln Met Val Val Phe Leu Leu Leu Val Ser Thr Leu Ser Ser
                        -10
                                            -5
Val Val Leu Leu Val Cys Ile Pro Thr Ser Ser Val Lys Leu Phe
                                    10
Pro Phe His His Ile His Thr Asn Trp
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<211> 30
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Met Glu Phe Gly Leu Ser Trp Val Leu Leu Val Ala Met Leu Arg Gly
               -15
                                    -10
Leu Gln Cys Gln Val Gln Leu Val Glu Ser Gly Gly Thr Ala
<210> 860
<211> 57
<212> PRT
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Met Tyr Leu Ser Leu Leu Ile Leu Leu Leu Glu Asn Val Ser Gly Phe
                   -10
                                       - 5
Pro Phe Pro Leu Ile Phe Gln Leu His Ala Ser Pro Gly His Lys Ile
         5
                               10
Leu Pro Asp Cys Met Ile Tyr Ser Ile Thr Val Ser Leu Met Phe Pro
                           25
Val Val Asp Tyr Ile Ser Thr Gln Gly
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<222> -28..-1
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Leu Trp Phe Xaa Cys Leu Leu Phe Leu Leu Phe Ala Trp Pro Gly
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<212> PRT
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                    -15
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Gly Ser Leu Ala Gln Leu Val Leu Thr Gln Ser Pro Ser Ala Ser Ala
Ser Leu Gly Ala Ser Val Lys Leu Thr Cys Thr Leu Ser Ser Gly His
Ser Asn Tyr Gly Ile Ala Trp Tyr Gln Gln Gln Pro Glu Lys Gly Pro
Arg Phe Leu Met Lys Val Asn Ser Asp Gly Ser His Met Lys Ala Asp
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Gly Ile Pro Asp Arg Phe Ser Gly Ser Ser Gly Ala Glu Arg Tyr
                65
Leu Ser Ile Ser Ser Leu
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                                    -5
Gly Gln
<210> 864
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<400> 864
Met Asp Trp Thr Trp Arg Xaa Phe Cys Leu Leu Ala Val Ala Pro Gly
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Ala His Ser Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys
Pro Gly Ala Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe
                        20
Thr Ser His Tyr Met His Trp Val Arg Gln Ala Pro Gly Gln Gly Leu
Glu Trp Met Gly Ile Ile Tyr Pro Asp Ser Asp Thr Thr Lys Tyr Xaa
Gin Asn Phe Gln Gly Arg Val Thr Met Thr Arg Asp Thr Ser Thr Ser
Thr Val Tyr Met Glu Leu Ser Ser Leu Thr Ser Asp Asp Thr Ala Val
                            85
Tyr Tyr Cys Ala Arg Glu Ala Tyr Ser Gly Ser Tyr Arg Phe Asp Tyr
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Trp 110 105

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 <221> SIGNAL
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Met Asp Leu Met Cys Lys Lys Met Arg His Leu Trp Phe Leu Leu Leu
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Leu Val Ala Ala Pro Arg Trp Val Leu Ser Gln Leu Gln Leu Gln Glu
                     -5
Ser Gly Pro Gly Leu Val Lys Ala Ser Glu Thr Leu Ser Leu Ala Cys
                                 15
Ser Val Ser Gly Asp Ser Ile Ser Ser Gly Asn Tyr Tyr Trp Gly Trp
                             30
 Ile Arg Gln Pro Pro Gly Lys Gly Leu Gln Trp Leu Gly Ser Leu Trp
                         45
Asn Arg Gly Gly Pro Gln Tyr Asn Xaa Ser Leu Lys Asn Arg Val Thr
                                         65
                     60
Val Ser Val Asp Thr Ser Thr Asn His Phe Phe Leu Arg Leu Asn Ser
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Val Asn Xaa Gly His Gly Asn Leu Leu Cys Ala
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 Met Arq Xaa Xaa Leu Xaa Leu Ser Val Leu Leu Gly Xaa Xaa Xaa
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                                          -5
 Lys Xaa Asp Phe Val Gly His Gln Val Leu Arg Ile Ser Val Ala Asp
                                     10
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 Met Ala Glu Ser Arg Glu Glu Gly Glu Ser Cys Val Glu Ser His Cys
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                                             -25
· Val Leu Phe Phe Thr Leu Phe Phe Leu Leu Phe Phe Cys Phe Val Phe
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 Cys Leu Arg Gly Gln Gly
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<212> PRT

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                                    -10
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Pro Gly Gly Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe
                       20
Ser Ser Tyr Ala Met Leu Trp Val Arg Gln Ala Pro Gly Lys Gly Leu
                   35
                                       40
Glu Trp Val Ser Gly Ile Ser Ala Gly Ala Asp Asp Thr Tyr Asp Ala
                                    55
Asp Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Lys
                               70
Ile Leu Tyr Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Arg
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           -20
                                                   -10
Leu Phe Leu Thr Cys Tyr Ala Asp Asp Lys Pro Asp Lys Pro Asp Asp
Lys Pro Asp Asp Ser Gly Lys Asp Pro Lys Pro Asp Phe Pro Lys Phe
                   15
                                       20
Leu Ser Leu Leu Gly Thr Glu Ile Ile Glu Asn Ala
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                                    -15
Ala Leu Pro Ser Pro Leu Ser Ala Arg Ala Glu Pro Pro Gln Asp Lys
Glu Ala Cys Val Gly Thr Asn Asn Gln Ser Tyr Ile Cys Asp Thr Gly
His Cys Cys Gly Gln Ser Gln Cys Cys Asn Tyr Tyr Tyr Glu Leu Trp
Trp Phe Trp Leu Val Trp Thr Ile Ile Ile Leu Ser Cys Cys
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Val Cys His His Arg Arg Ala Lys His Arg Leu Gln Ala Gln Gln Arg
60 65 70

454 Gln His Glu Ile Asn Leu Ile Ala Tyr Arg <210> 871 <211> 37 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -27..-1 <400> 871 Met Val Val Ala Asp Arg Asn Arg Ala Ser Ser Ser Tyr Leu Cys -20 Leu Leu Phe Ser Leu Ser Leu Phe Leu Cys His Glu Thr Val Cys Asp Arg Ala Thr Cys <210> 872 <211> 142 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -19..-1 <400> 872 Met Asp Trp Thr Trp Arg Phe Leu Phe Val Val Ala Ala Ala Thr Gly -15 -10 Val Gln Ser Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Ser Ser Val Lys Val Ser Cys Lys Ala Ser Gly Gly Thr Phe 20 Ser Xaa Tyr Ala Ile Ser Trp Val Arg Gln Ala Pro Gly Gln Gly Leu Glu Trp Met Gly Gly Ile Ile Pro Ile Phe Gly Thr Ala Xaa Tyr Ala 50 55 Gln Lys Phe Gln Gly Arg Val Thr Ile Thr Ala Asp Xaa Ser Thr Xaa 70 Thr Xaa Tyr Met Glu Leu Ser Ser Leu Arg Ser Glu Asp Thr Ala Xaa 85 Tyr Tyr Cys Ala Arg Gly Gln Ala Pro Gly Arg Val Val Pro Leu 100 105 Phe Leu Trp Gly Gln Gly Thr Trp Ser Pro Ser Pro Gln Pro 115 <210> 873 <211> 87 <212> PRT <213> Homo sapiens <220> <221> SIGNAL * 25.--<222> -45..-1 <400> 873 Met Thr Tyr Ser Tyr Ser Phe Phe Arg Pro Glu Leu Ile Val Asn His -40 -35

Leu Asn Tyr Val His Ser Glu Ala Asn Arg Arg Thr Lys Thr Lys Thr

-20

-25

455

Leu Leu Ser Leu Leu Ser Phe Leu Asp Glu Thr Ser Gly Leu Ser Thr -10 -5 His Leu Pro Cys Leu Ser Leu Ser Lys Glu Cys Gly Val Leu His Leu 10 Asp Ile His Gly Lys Lys Glu Asp Met Arg Asp Glu Val Leu Leu Ala 25 Leu Asn Xaa Cys Thr His Arg 40 <210> 874 <211> 79 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -19..-1 <400> 874 Met Lys Ser Phe Ser Arg Ile Leu Phe Leu Val Phe Leu Leu Ala Gly -15 -10 Leu Arg Ser Lys Ala Ala Pro Ser Ala Pro Leu Pro Leu Gly Cys Gly 5 1 Phe Pro Asp Met Ala His Pro Ser Glu Thr Ser Pro Leu Lys Gly Ala 20 Ser Glu Asn Ser Lys Arg Asp Arg Leu Asn Pro Glu Phe Pro Gly Thr 35 40 Pro Tyr Pro Glu Pro Ser Lys Leu Pro His Thr Val Ser Leu Glu <210> 875 <211> 51 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -41..-1 <400> 875 Met Arg Val Pro Ile Phe Pro His Pro His Gln Leu Ser Leu Leu Phe -35 -30 Ile His Leu Phe Ile Tyr Leu Phe Arg Glu Arg Val Ser Leu Cys His -20 -15 Leu Gly Trp Ser Ala Val Val Gln Ser Gln Pro Thr Thr Leu Thr Ser Arg Ala 10 <210> 876 <211> 44 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -37..-1 <400> 876 Met Trp Lys Glu Ser Ser His Gly Cys Asn Asn Leu Gly Ser Ser Tyr -25 -30 -35 Leu Asp Asp Thr Gly Val Gly Ser Phe Leu Phe Val Leu Phe Cys Phe

-15

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Gly Gly Ser Arg Ala Leu Leu Leu Pro Gly Ser Gly
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                       -10
Ala Val Phe Leu Thr Val Pro Ser Pro Gln
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<221> SIGNAL
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                -35
Xaa Arg Xaa Ser Leu Pro Ala Cys Ala Asp Ser Ile Ile Leu Xaa Leu
                                -15
           -20
Xaa Phe Pro Gly Ile Leu Gly Gln Ala His Leu Xaa Ser Glu Gln Trp
        -5
Thr Gln Tyr Leu
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                                    -10
 Gly Thr Cys Phe Thr Trp Ile Leu Leu Trp Leu Pro Leu Ser Pro Leu
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                     1
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 Leu Gly Leu Lys Cys
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 ₹233~ 85 °
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70

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459 Glu Trp Leu Gly Arg Ile Ile Pro Ile Leu Gly Ile Thr Asn Tyr Ala 55 60 Glu Lys Phe Arg Gly Arg Leu Thr Ile Thr Val Asp Lys Ser Thr Arg Val Val Tyr Met Glu Gln Ser Ser Leu Thr Ser Ala Asp Thr Ala Val 85 Tyr Tyr Cys Ala Lys Pro Thr Met Thr Ser Glu Leu Arg Val Tyr Tyr 100 Gln Xaa Thr Leu Trp <210> 886 <211> 30 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -22..-1 <400> 886 Met Trp Asn Arg Tyr Phe. Val Phe Tyr Leu Leu Leu Ser Ala Phe -20 -15 Thr Ser Gln Thr Val Ser Gly Gln Arg Lys Lys Gly Pro Arg <210> 887 <211> 142 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -19..-1 <400> 887 Met Lys His Leu Gly Phe Phe Leu Leu Leu Val Ala Ala Pro Arg Trp -10 . -15 Val Leu Ser Gln Leu Gln Leu Gln Glu Ser Gly Ser Gly Leu Glu Lys Pro Ser Gln Thr Leu Ser Leu Thr Cys Ser Val Ser Gly Gly Ser Ile 20 Ser Ser Asp Asp Leu Ser Trp Ser Trp Ile Arg Gln Pro Pro Gly Lys 40 35 Gly Leu Glu Trp Ile Gly Tyr Ile Tyr Gln Asn Glu Arg Thr Leu Tyr 55 50 Asn Pro Ser Leu Lys Ser Arg Ala Ala Ile Ser Val Asp Arg Ser Lys 70 Asn Gln Phe Ser Leu Lys Leu Thr Ser Val Thr Ala Ala Asp Met Ala 85 Val Tyr Tyr Cys Ala Thr Ser Val Met Xaa Ser Phe Gly Gly Val Leu Val Pro Asn Leu Phe Leu Thr Thr Gly Ala Arg Glu Ser Arg 115

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<212> PRT

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<220>

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                                   -10
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Pro Ser Gln Thr Leu Ser Leu Thr Cys Thr Val Ser Gly Ala Ser Val
Ser Ser Arg Gly Tyr Tyr Trp Thr Trp Ile Arg Gln Leu Pro Gly Lys
                    35
Gly Leu Glu Trp Ile Gly Tyr Ile Xaa Tyr Thr Gly Ser Thr Phe Tyr
                                    55
Asn Pro Ser Leu Lys Ser Arg Leu Thr Ile Ser Ile Asp Thr Ser Lys
                                70
Asn Gln Phe Ser Leu Asn Leu Arg Ser Val Thr Thr Ala Asp Thr Ala
                           85
Val Tyr Tyr Cys Ala Arg Asp His Phe Asp Leu Leu Phe Asp Pro Trp
                                           105
                        100
Gly Gln Gly Thr Leu Val Thr Val Ser Ser Ala Ser Thr Lys Gly Pro
                                       120
                115
Ser Val Phe Pro Leu Ala Xaa Ser Ser Lys Ser
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                        -35
                                            -30
Arg Phe Phe Asn Trp Gly Lys Leu Phe Phe Cys Phe Val Leu Xaa Leu
                                     -15
                    -20
Phe Cys Phe Val Phe Glu Ala Glu Ser Arg Ser Val Ala Gln Ala Gly
                                 1 .
                -5
Val Gln Trp Arg Tyr Phe Gly Ser Leu Gln Ala Leu Pro Pro Trp
                            15
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 Met His Glu Phe Ile Ser Gly Phe Phe Ile Leu Phe His Trp Ser Leu
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 Cys Leu Cys Leu Cys Gln Tyr His Ala
 <210> 891
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461

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Met Ala Tyr Ala Ile Ser Pro Phe His Ser Ser Trp Asn Pro Leu Phe
                  -35
Thr Ser His Lys Ala Ser Ala Ser His Ser His Leu Gly Leu Leu Val
                      -20
Cys Leu Phe Ala Val Thr Ser Ile Leu Cys Ser Ser
<210> 892
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                                        ~ 5
                   -10
Pro Gly Ser Ala Xaa Ala Xaa Ser Ala Ser Leu Gly Gln Phe Ser Met
                               10
Cys Gly Arg Cys Pro Thr Cys Pro Gly Asn Gly Pro Leu Arg Thr Pro
                           25
Ala Ala Thr Xaa Xaa Xaa Val Pro Gly His Val Asp
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<210> 893
<211> 154
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<400> 893
Met Ala Thr Ala Met Asp Trp Leu Pro Trp Ser Leu Leu Leu Phe Ser
                                ~15
Leu Met Cys Glu Thr Ser Ala Phe Tyr Val Pro Gly Val Ala Pro Ile
Asn Phe His Gln Asn Asp Pro Val Glu Ile Lys Ala Val Lys Leu Thr
                                        20
                    15
Ser Ser Arg Thr Gln Leu Pro Tyr Glu Tyr Tyr Ser Leu Pro Phe Cys
                                    35
                30
Gln Pro Ser Lys Ile Thr Tyr Lys Ala Glu Asn Leu Gly Glu Val Leu
                                50
            45
Arg Gly Asp Arg Ile Val Asn Thr Pro Phe Gln Val Leu Met Asn Ser
                            65
Glu Lys Lys Cys Glu Val Leu Cys Ser Gln Ser Asn Lys Pro Val Thr
                                            85
                        80
Leu Thr Val Glu Gln Ser Arg Leu Val Ala Glu Arg Ile Thr Glu Asp
                                        100
                    95
Tyr Tyr Val His Leu Ile Ala Asp Asn Leu Pro Val Ala Thr Gly Trp
                                   115
                110
Ser Ser Thr Pro Thr Glu Thr Ala Met Thr
            125
 <210> 894
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<211> 28

462

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<212> PRT
<213> Homo sapiens
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Met Pro Ser Pro Cys Leu Ile Ser Leu Leu Gln Cys Ala His Val Ser
                                -10
           -15
Leu Gly Leu Gln Tyr Pro Cys Xaa Leu Leu Leu Pro
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<210> 895
<211> 53
<212> PRT
<213> Homo sapiens
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<222> -17..-1
<400> 895
Met Asn Leu Ser Leu Val Leu Ala Ala Phe Cys Leu Gly Ile Ala Ser
                             -10
      -15
Ala Val Pro Lys Phe Asp Gln Asn Leu Asp Thr Lys Trp Tyr Gln Trp
                    5
                                         10
Lys Ala Thr His Arg Arg Leu Tyr Gly Ala Asn Glu Glu Gly Trp Arg
                                     25
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Arg Ala Ala Trp Glu
            35
<210> 896
<211> 85
 <212> PRT
<213> Homo sapiens
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 <222> -19..-1
 <400> 896
Met Glu Phe Gly Leu Asn Trp Val Phe Leu Val Ala Ile Phe Thr Gly
                                     -10
                 -15
 Val His Cys Glu Val Gln Leu Val Glu Ser Gly Gly Asp Leu Val Gln
                                                  10
 Pro Gly Arg Ser Leu Arg Leu Ser, Cys Thr Ala Ser Gly Phe Thr Phe
                                              25
                         20
 Gly Asp Tyr Ala Met Thr Trp Phe Arg Gln Ala Ser Gly Lys Arg Leu
                                          40
                     35
 Glu Trp Leu Gly Phe Ile Arg Asn Arg Gly Ser Gly Gly Ser Ala Glu
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 Tyr Gly Ala Ser Val
             65
 <210> 397
 <211> 51
 <212> PRT
 <213> Homo sapiens
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<222> -17..-1

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Met Lys Asn Cys Leu Leu Ile Leu Leu Met Leu Leu Phe Ala Ile
       -15
                           -10
His Ile Asn Arg Met Asn Val Arg Asn Val Gly Asn Thr Leu Val Val
                5
                                       10
Val Gln Ile Leu Phe Ser Ile Arg Val Phe Ile Leu Glu Arg Asn Pro
Leu Asn Val
<210> 898
<211> 149
<212> PRT
<213> Homo sapiens
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<221> SIGNAL
<222> -19..-1
<400> 898
Met Glu Leu Gly Leu Ser Trp Ile Phe Leu Leu Ala Ile Leu Lys Gly
               -15
                          -10
Val Gln Cys Glu Val Gln Leu Val Glu Ser Gly Gly Leu Val Gln
                        5
Pro Gly Arg Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe
                      20
Asp Asp Tyr Ala Met His Trp Val Arg Gln Ala Pro Gly Lys Gly Leu
                35
Glu Trp Val Ser Gly Ile Thr Trp Asn Ser Gly Xaa Ile Gly Tyr Ala
              50
                                  55
Asp Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn
        65
                              70
Ser Leu Tyr Leu Gln Met Asn Ser Leu Arg Thr Glu Asp Thr Ala Phe
                          85
Tyr Phe Cys Ala Lys Ala Arg Gly Leu Phe Ser Asp Thr Trp Pro Tyr
                      100
                                          105
Xaa His Tyr Ala Met Asp Val Trp Gly Gln Gly Thr Thr Val Thr Val
                   115
                                       120
Ser Ser Ala Ser Thr
<210> 899
<211> 25
<212> PRT
<213> Homo sapiens
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Met Leu Leu Val Phe Phe Val Leu Trp Thr Cys Ser Leu Ala Leu Leu
               -10
Ala Ser Ser Pro Ile Ala Ala Xaa Pro
       5
                           10
<210> 900
<211> 127
<212> PRT
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<220>
<221> SIGNAL
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<222> -19..-1
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-35 Asp Pro Arg Ala Tyr Phe Lys Thr Lys Thr Trp Trp Leu Gly Leu Phe -20

Leu Met Leu Gly Glu Leu Gly Val Phe Ala Ser Tyr Ala Phe Ala

-15

Pro Leu Ser Leu Ile Val Pro Leu Ser

<210> 903 <211> 44 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -18..-1 <400> 903 Met Ala Phe Leu Trp Leu Leu Ser Cys Trp Ala Leu Leu Gly Thr Thr -15 -10 Phe Gly Cys Gly Val Pro Ala Ile His Pro Gly Cys Gln Leu Ser Pro 5 1 10 Arg Leu Pro Pro Thr Leu Leu Pro Thr Glu Arg Gly 20 <210> 904 <211> 82 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -20..-1 <400> 904 Met Ala Pro Phe Gln Asn Phe Leu Trp Leu Phe Phe Val Leu Asn Leu -15 -10 Gly Ser Phe Ala Phe Ser Ser Xaa Pro Asn Ser Leu Phe Tyr Thr Ile 1 10 His Phe Gly Pro Asn Phe Phe Thr Leu Leu Tyr Lys Gln Gly Ala Glu 20 Met Cys Val Tyr Val Phe Asn Phe Leu Tyr Pro Phe Ala Leu Gly Tyr 35 40 Phe Phe Ser Tyr Asp Ile Leu Asp Leu Pro Val Xaa Val Arg Pro Pro 50 Ser Gly <210> 905 <211> 54 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -35..-1 <400> 905 Met Asp Phe Thr Gln Cys His Ser Leu Leu Leu Arg Val Glu Tyr Ser -30 - 25 Pro Val Ser Val Cys Phe Leu Leu Ser Val Ala Phe Asn Gln Leu -15 -10 Val Phe Ala Leu Tyr Pro Ile Gln Ala Thr Xaa Cys Phe Ser Xaa Val

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<210> 906 <211> 23

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Ser Leu Pro Phe Pro Ala

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<212> PRT
<213> Homo sapiens
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<221> SIGNAL
<222> -15..-1
<400> 906
Met Leu Leu Leu Leu Ala Cys Gly Val Pro Ser Leu Trp Pro Phe
                   -10
                                       -5
Ala Leu Ala Leu Leu Lys Thr
           5
<210> 907
<211> 43
<212> PRT
<213> Homo sapiens
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<221> SIGNAL
<222> -23..-1
<400> 907
Met Phe Ile Glu Asn Ile Gly Leu Lys Phe Ser Phe Leu Leu His
          -20
                              -15
Leu Cys Gln Val Leu Leu Ser Arg Arg Ala Gly Thr Ile Pro Thr Glu
 -5
                          1
Thr Ile Pro Lys Lys Leu Arg Arg Arg Asp Gly
                  15
<210> 908
<211> 105
<212> PRT
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<222> -24..-1
<400> 908
Met Gln Asn Arg Thr Gly Leu Ile Leu Cys Ala Xaa Ala Leu Leu Met
               -20
                                   -15
Gly Phe Leu Met Val Cys Leu Gly Ala Phe Phe Ile Ser Trp Gly Ser
                               1
Ile Phe Asp Cys Gln Gly Ser Leu Ile Ala Ala Tyr Leu Leu Pro
Leu Gly Phe Val Ile Leu Leu Ser Gly Ile Phe Trp Ser Asn Tyr Arg
Gln Val Thr Glu Ser Lys Gly Val Leu Arg His Met Leu Arg Gln His
               45
                                   50
Leu Ala His Gly Ala Leu Pro Val Ala Thr Val Asp Ser Ala Ala Leu
           60
Leu Lys Ile Met Cys Lys Gln Leu Leu
<210> 909
<211> 52
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
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<222> -44..-1

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Met Lys Val Glu Glu Glu Lys Leu Tyr Arg Leu Leu Arg Ser Gly
               -40
                                   -35
Asp Leu Phe Lys Phe His Gln Pro His Phe Tyr Glu Leu Ser Gly Leu
                             -20
         -25
Thr Cys Thr Ser Ser Leu Leu Ser Phe Ala Leu Gly Arg Ser Ile Pro
 -10
                        -5
Gly Ser Phe Pro
<210> 910
<211> 60
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -19..-1
<400> 910
Met Glu Ser Arg Thr Leu Leu Leu Phe Ser Gly Ala Val Ala Leu
               -15
                                   -10
Ile Gln Thr Trp Ala Gly Glu Cys Gly Val Gly Arg Glu Lys Ala Ser
Ala Gly Arg Ser Glu Gly Pro Ala Arg Arg Ser Lys Ser Ala His Ile
                       20
                                           25
Xaa Asn Tyr Arg Leu Gln Leu Gln Ser Arg Gln Gly
                   35
<210> 911
<211> 35
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -16..-1
<400> 911
Met Ser Asn Ser Val Pro Leu Cys Phe Trp Ser Leu Cys Tyr Cys
                                           -5
                       -10
Phe Ala Ala Gly Ser Pro Val Pro Phe Gly Pro Glu Gly Arg Leu Glu
                                   10
Asp Lys Leu
<210> 912
<211> 52
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -14..-1
<400> 912
Met Pro Trp Thr Ile Leu Leu Phe Ala Ala Gly Ser Leu Ala 11e Fro
               -10
                            -5
Ala Pro Ser Ile Arg Val Val Pro Pro Tyr Pro Ser Ser Gln Glu Asp
                          10
Pro Ile His Ile Ala Cys Met Ala Ala Gly Asn Phe Pro Gly Ala Asn
   20
                       25
Phe Thr Leu Tyr
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<210> 913 <211> 67 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -64..-1 <400> 913 Met Ala Glu Gly Glu Arg Val Cys Ala Ser Val Val Pro Ser Ala Leu -60 -55 Arg Thr Leu Lys Arg Arg Ser Asn Leu Ser Arg Ile Pro Ala Gly Gln -40 -45 -35 Glu Lys Glu Gly Lys Ser Arg His Val Ala Pro Pro Phe Arg Phe Phe -25 -20 Pro Phe Ser Gly Phe Leu Phe Phe Gly Phe Leu Phe Pro Val Phe Ser -15 -10 Phe Pro Ser <210> 914 <211> 71 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -13..-1 <400> 914 Met Phe Cys Leu Ala Ala Ile Leu Ala Ser Ala Ser Ala Gln Arg Phe -10 -5 Pro Ser Ala Phe Ser Pro Ser Pro Phe Xaa Trp Leu Xaa Gln Cys Xaa 10 Thr Ala Thr Ser Leu Gly Phe Xaa Thr Val Cys Xaa Asn Ser Ile Ile 25 30 Ser Leu Trp Tyr Leu Xaa Gly Val Pro Pro Glu Val Xaa Glu Leu Pro 40 45 Phe Phe Pro Tyr Cys Ser Met 55 <210> 915 <211> 93 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -17..-1 <400> 915 Met Val Asp Gly Thr Leu Leu Leu Leu Ser Glu Ala Leu Ala Leu -15 -10 -5 Thr Gln Thr Trp Ala Gly Ser His Ser Kaa Lys Tyr Phe His Thr Cor 15 10

Val Ser Arg Xaa Gly Arg Gly Glu Pro Arg Phe Ile Ser Val Gly Tyr

Val Asp Asp Thr Arg Ser Glu Tyr Trp Asp Arg Glu Thr Arg Ser Ala 40 Arg Asp Thr Ala Gln Ile Phe Arg Val Asn Leu Arg Thr Leu Arg Gly

25

20

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469
       50 ·
Tyr Tyr Asn Gln Ser Glu Ala Gly Ser Xaa Thr Leu Gln
                        70
<210> 916
<211> 75
<212> PRT
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Met Asn Phe Arg Gly Pro Gln Thr Phe Ser Leu Ser His Ser Leu Val
       -25
                            -20
                                                 -15
Leu Ser Leu Ile Ser Leu Ser Ile Ala Trp Ser Met Val Glu Met Xaa
   -10
                        -5
Thr Ser Ala Ser Tyr Lys Gln Lys Phe Ala Leu Arg Ile Leu Val Val
                10
                                    15
Gln Leu Pro Thr Trp Val Glu Cys Pro Val Asn His Arg Cys Ala Leu
                                30
                                                     35
Gly Arg Lys Asn Cys Ser Ile Arg Thr Gln Pro
<210> 917
<211> 25
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<213> Homo sapiens
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<221> SIGNAL
<222> -20..-1
<400> 917
Met Thr Gly Ile Ser Ile Cys Ser Cys Ile Cys Leu Phe Leu Pro Ser
                    -15
                                         -10
Leu Ile His Ser Phe Pro Pro Cys
                1
<210> 918
<211> 98
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -26..-1
<400> 918
Met Asp Leu Cys Lys Asn Met Lys His Leu Trp Phe Phe Leu Leu
   -25
                        -20
                                            -15
Leu Val Ala Ala Pro Arg Trp Val Gln Leu Gln Glu Ser Gly Pro Arg
                    -5
Leu Val Arg Pro Pro Glu Thr Leu Lys Pro Ser Glu Thr Leu Ser Leu
                                15
Thr Cys Thr Ile Ser Gly Asp Ser Met Ser Ser Ala Ser Tyr Tyr Trp
Ala Trp Ile Arg Gln Pro Pro Gly Lys Gly Leu Glu Phe Ile Gly Arg
                        45
Ala Leu Tyr Ser Gly Thr Thr Asp Tyr Asn Pro Ser Leu Ser Ser Arg
55
Ile Thr
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<210> 919
<211> 52
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -45..-1
<400> 919
Met Ser Ser Glu Lys Ser Gly Leu Pro Asp Ser Val Pro His Thr Ser
                   -40
                                        -35
Pro Pro Pro Tyr Asn Ala Pro Gln Pro Pro Ala Glu Pro Pro Ala Pro
                -25
                                    -20
Pro Leu Ser Leu Cys Leu Ser Leu Cys His Thr His Thr His
                                -5
Thr His Thr His
    5
<210> 920
<211> 46
<212> PRT
<213> Homo sapiens
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<221> SIGNAL
<222> -28..-1
<400> 920
Met Thr Pro Ala Leu Arg Cys Ala Phe Ala Leu Ala Ile Ala Gly Leu
           -25
                                -20
                                                    -15
Val Ser Leu Leu Met Gln Pro Glu Gly Ala Leu Gly Glu Glu Ala Ala
                            -5
Ser Ala Ala Ala Gln Gly Arg Gln Leu Ala Glu Leu Arg Leu
                    10
<210> 921
<211> 70
<212> PRT
<213> Homo sapiens
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<221> SIGNAL
<222> -38..-1
<400> 921
Met Ser Gly Leu Phe Pro Val Pro Val Arg Val Asn Val Asp Ile Ala
            -35
                                -30
                                                    -25
Gln Asn Ile Thr Cys Ser Ser Phe Ser Leu Leu Leu Ile Phe Leu Ser
                            -15
                                                -10
Phe Pro Tyr Thr Leu Cys Ile Leu Tyr Arg Val Lys Ser Tyr Thr Pro
                        1
                                        5
Thr Glu Ser Ile Thr Ala Phe Asn Leu Thr Ile Glv Xaa Phe Pro Tyr
                15
                                    20
Leu Xaa Xaa Ser Thr Pro
            30
<210> 922 ·
<211> 39
<212> PRT
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471 <220> <221> SIGNAL <222> -33..-1 <400> 922 Met Cys Arg Ala Ala Cys Ile Ile Arg Met Ala Val Arg Ile Ser Phe -25 Phe Leu Ser Tyr His Ala Leu Ser Leu Cys Leu Cys Thr Cys Ala Phe -10 Ala Phe Leu Ser Leu Leu Gly 1 <210> 923 <211> 59 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> ~17..-1 <400> 923 Met Lys Phe Leu Leu Leu Xaa Ala Leu Gly Phe Leu Xaa Gln Val Asn -15 ~10 Pro Xaa Pro Ile Xaa Gly Gly Ser Lys Met Cys Glu Xaa His Pro Arg 10 Ile Leu Gln Asp Met Leu Pro Leu Gly Gly Asp Ser Ile Val His Val 20 Gln Arg Xaa Gln Lys Met Leu His Gln Leu Leu <210> 924 <211> 105 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -42..-1 <400> 924 Met Val Pro Trp Val Arg Thr Met Gly Gln Lys Leu Lys Gln Arg Leu -35 Arg Leu Asp Val Gly Arg Glu Ile Cys Arg Gln Tyr Pro Leu Phe Cys -20 -15 Phe Leu Leu Cys Leu Ser Ala Ala Ser Leu Leu Leu Asn Arg Tyr -5 Ile His Ile Leu Met Ile Phe Trp Ser Phe Val Ala Gly Val Val Thr 10 15 Phe Tyr Cys Ser Leu Gly Pro Asp Ser Leu Leu Pro Asn Ile Phe Phe 30 Thr Ile Lys Tyr Lys Pro Lys Gln Leu Gly Leu Gln Glu Leu Phe Pro 45 Gln Gly His Ser Cys Ala Val Cys Gly 60 <210> 925 <211> 43 <212> PRT <213> Homo sapiens <220>

<221> SIGNAL

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Met Ala Trp Gly Ser Pro Gly Lys Ile Phe Leu Met Gly Phe Leu Gly
               -30
                                   -25
Gly Glu Leu Val Phe Leu Leu Cys Leu Phe Xaa Leu Phe Phe Phe Ser
           -15
                               -10
Phe Leu Lys Arg Ser Phe Ala Leu Glu Cys Asn
                       5
<210> 926
<211> 28
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -16..-1
<400> 926
Met Phe Phe Ser Ile Leu Leu Leu Ala Pro Pro Leu Pro Ser Ala
                        -10
                                           -5
Val Ser Leu Leu Pro Phe Phe Phe Tyr Cys Val Gln
               5
                                   10
<210> 927
<211> 42
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -22..-1
<400> 927
Met Val Asp Phe Ile Leu Arg Ser Leu Leu Leu Val Cys Ser Trp Leu
                           -15
                                               -10
Ser Ile Ser Leu His Ala His Thr Thr Ala Phe Cys Thr Tyr Ser Lys
                . 1
                                                           10
Lys Ile His Thr Val Met Ser Phe Phe Cys
               15
<210> 928
<211> 26
<212> PRT
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<222> -16..-1
<400> 928
Met Arg Ser Leu Leu Tyr Phe Leu Cys Val Ser Ser Tyr Val Thr Ser
           _-10
Phe Phe Phe Phe Phe Phe Phe Phe
<210> 929
<211> 68
<212> PRT
<213> Homo sapiens
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<220>

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473
<221> SIGNAL
<222> -15..-1
<400> 929
Met Pro Phe Ile Ser Phe Leu Cys Leu Ile Ala Leu Ala Gly Thr Ser
                   -10
Ser Thr Met Leu Arg Ser Ala Leu Ala Gly Thr Ser Ser Thr Met Xaa
                               10
                                                  15
Xaa Arg Ser Gly Xaa Ser Gly Xaa Pro Xaa Leu Val Xaa Val Leu Arg
                          25
                                             30
Gly Asn Ala Phe Ser Phe Phe Pro Phe Ser Leu Met Xaa Ala Met Gly
  35
                       40
Cys His Arg Trp
50
<210> 930
<211> 22
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<222> -16..-1
<400> 930
Met Tyr Thr Phe Leu Leu Gly Ala Ile Phe Ile Ala Leu Ser Ser Ser
-15 -10
Arg Ile Leu Leu Val Lys
<210> 931
<211> 44
<212> PRT
<213> Homo sapiens
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<221> SIGNAL
<222> -42..-1
<400> 931
Met Cys Leu Cys Pro Cys Trp Asp Val Phe Thr Val Phe Val Cys Val
  -40
             · -35
Ser Val Cys Val Ser Val Ser Val Pro Val Gly Met Tyr Leu Val Cys
                       -20
Val Cys Val Cys Val Cys Xaa Cys Xaa Arg
<210> 932
<211> 50
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -34..-1
<400> 932
Met Leu Ile Ala Lys Gln Ala Gln Pro Gln Gly Leu Thr Ala Ile Cys
                                   -25
               -30
Phe Pro Leu Thr Pro Leu Phe Ser Leu Leu Met Leu Thr Gln Ser Pro
                               -10
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Leu Ala Gly Gln Glu Gly Arg Glu Gly Gly Lys Glu Arg Tyr Leu Leu

10

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WO 99/53051
                                      474
Val Ile
15
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<211> 62
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<213> Homo sapiens
<220>
<221> SIGNAL
<222> -26..-1
<400> 933
Met Leu Arg Thr Trp Ser Ser Leu Pro Trp Thr Arg Phe Arg Val Cys
                        -20
                                            -15
Leu Leu Ser Leu Ser Leu Phe Leu Trp Ala Asn Arg Leu Glu Asp Ser
                    -5
                                        1
Arg Ser Cys Gln Pro Asn Pro Met Ser Leu Thr Thr Leu Pro Gly His
          10
                               15
Arg Leu Lys Glu Ala Val Trp Leu Pro Ala Pro Ser Leu Gly
                           30
<210> 934
<211> 72
<212> PRT
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<400> 934
Met Ala Pro Phe Leu Arg Gln Val Asp Xaa Trp Gly Ala Gln Ala Gly
                -25
                                    -20
Leu Val Val Xaa Trp Leu Leu Pro Xaa Gln Cys Ser Cys Glu Arg Ser
           -10
                                -5
Glu Gln Tyr Leu Ser Thr Cys Leu Pro Gln His Ser Ser Ile Lys Gln
                        10
Ser Cys Ile Lys His Pro Ala Gly Pro Ile Pro Ala Gly His Leu Gln
                   25
Gly Lys Ala Thr Ala Ala Pro Leu
<210> 935
<211> 73
<212> PRT
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<222> -19..-1
<400> 935
Met Glu Phe Gly Leu Lys Trp Leu Phe Leu Val Ala Ile Leu Lys Gly
                -15
                                    -10
Val Arg Cys Glu Val Lys Leu Val Glu Ser Gly Gly Leu Val Gln
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Pro Gly Gly Ser Leu Arg Leu Ser Cys Val Gly Ser Gly Phe Val Phe Asp Lys Tyr Gly Ile Ser Trp Val Arg Gln Ala Pro Gly Lys Gly Leu 35 Gln Trp Val Ala Gly Ile Gly Gly Gly 50

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<211> 128
<212> PRT
<213> Homo sapiens
<220>
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<222> -16..-1
<400> 936
Met Ala Leu Ala Met Leu Val Leu Val Val Ser Pro Trp Ser Ala Ala
                       -10
Arg Gly Val Leu Arg Asn Tyr Trp Glu Arg Leu Leu Arg Lys Leu Pro
                                   10
Gln Ser Arg Pro Gly Phe Pro Ser Pro Pro Trp Gly Pro Ala Leu Ala
                               25
Val Gln Gly Pro Ala Met Phe Thr Glu Pro Ala Asn Asp Thr Ser Gly
                           40
                                               45
Ser Lys Glu Asn Ser Ser Leu Leu Asp Ser Ile Phe Trp Met Ala Ala
                      55
Pro Lys Asn Arg Arg Thr Ile Glu Val Asn Arg Cys Arg Arg Arg Asn
                   70
                                       75
Pro Gln Lys Leu Ile Lys Val Lys Asn Asn Ile Asp Val Cys Pro Glu
               85
                                  90
Cys Gly His Leu Lys Gln Lys Xaa Val Leu Cys Ala Thr Ala Met Lys
                              105
<210> 937
<211> 30
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<213> Homo sapiens
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<221> SIGNAL
<222> -20..-1
<400> 937
Met Phe Phe Tyr Ser His Phe Leu Leu Phe Pro Leu Ser Leu Leu
                  -15
                                       -10
Phe Thr Leu Gly Phe Leu Phe Val Phe Phe Phe Phe Phe
               1
                    5
<210> 938
<211> 101
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -46..-1
<400> 938
Met Lys Gln Ser Lys Arg Xaa Met Val Lys Arg Arg Arg Ser Pro Ala
                       -40
                                            35
Leu Gly Glu Glu Arg Phe Ser Pro Ser Ser Ile Leu His Pro Arg Leu
                   -25
                                       -20
Pro Leu Val Leu Leu Gly Thr Arg Val Pro Leu Ser Gly Gly Pro
                                   -5
Gly Glu Pro Asp Gln Gly Arg Ser Ala Pro Ser Trp Lys Ser Leu Ala
       5 -
                           10
                                               15
Ser Thr His Xaa His Ser Arg Pro Ala Ala Gly Ala Thr Pro Ala Arg
```

20

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Pro Ala Thr Gln Ser Gln-Leu Gly Pro Phe Ala Pro Pro Leu Pro Gly
                    40
                                        45
Val Arg Pro Ala Pro
<210> 939
<211> 32
<212> PRT
<213> Homo sapiens
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<222> -18..-1
<400> 939
Met Leu Leu Glu Ser Leu Cys Val Leu Ser Leu Leu Val Ser Phe Lys
                               -10
          -15
Ser Ala Cys Leu Thr Arg Glu Pro Ala Phe Asp Ser Gln Ala Arg Pro
<210> 940
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<400> 940
Met Val Phe Gly Tyr Trp Lys Gln Pro Leu Ile Thr Leu Ala Lys Lys
                        -40
                                            -35
Ser Val Lys Cys Ala Arg Glu Cys Leu Arg Cys Ser Leu Arg Pro Leu
                   -25
                                        -20
Val Leu Leu Tyr Leu Ser Phe Ala Ala Leu Gly Val Val Ala Leu Arg
                -10
                                   -5
Ser Val Glu Ser Pro Leu Ala Glu Thr His Ser Cys Trp Leu Ser Leu
                           10
                                               15
Gly Met Cys Val Leu Gln Cys Glu Gln Gln Trp Val Pro Thr Pro Val
                       25
Ser Phe Leu Cys Gly Leu Ser Gly Ser Ser Thr Ile Ile Val
<210> 941
<211> 66
<212> PRT
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<222> -24..-1
<400> 941
Met Cys Val Val Cys Ser Val His Gly Val Cys Cys Val Tyr Val Val
                                    -15
                -20
Cys Leu Val Ser Cys Val Leu Cys Val Val Cys Pro Val Cys Trp Val
Met Cys Cys Val Trp Cys Ile Cys Val Cys Val Trp Cys Val Cys Cys
                        15
                                            20
Met Cys Cys Val Leu Ser Cys Val Val Ser His Gly Leu Cys Gly Val
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25

Ser Trp

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477
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Met Glu Leu Gly Leu Ser Trp Val Phe Leu Val Ala Val Leu Glu Val
                -15
                                    -10
Val Gln Cys Glu Ile Gln Leu Ile Asp Ala Gly Gly His Val Gln
                            5
                                                 10
Ala Gly Gly Ser Leu Arg Leu Ser Cys Val Ala Ser Asp Phe Leu Phe
                        20
Arg Ser Tyr Trp Met Thr Trp Val Arg His Pro
                    35
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<211> 41
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<222> -39..-1
<400> 943
Met Ser Ile Leu Leu Arg Val Leu Gly Ile Lys Gly Cys Trp Ile Leu
                -35
                                    -30
                                                        -25
Ser Asn Pro Phe Ser Ala Cys Ile Glu Met Ile Leu Leu Phe Leu Phe
            -20
                                -15
                                                     -10
Leu Ile Leu Phe Ile Trp His Ile Arg
       ~5
<210> 944
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Met Ala Glu Lys Ala Gly Ser Thr Phe Ser His Leu Leu Val Pro Ile
                    -20
                                        -15
Leu Leu Ile Gly Trp Ile Val Gly Cys Thr
                -5
<210> 945
<211> 34
<212> PRT
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<220>
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<222> -19..-1
<400> 945
Met Ala Glu Ser Arg Gly Arg Leu Tyr Leu Trp Met Cys Leu Ala Ala
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-15

Ala Leu Ala Ser Phe Leu Met Gly Phe Met Val Gly Trp Phe Ile Lys

Pro Leu 15

<210> 946

<211> 40

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -26..-1

<400> 946

Met Leu Thr Ser Leu Pro Phe Leu Leu Pro Thr Ile Ser Phe Leu Leu
-25 -20 -15

Leu Leu Tyr Phe Phe Xaa Ile Ala Val Thr His Pro Ser Val Leu Ile
-10 -5 1 5

Asn Phe Ser Phe Ser Phe Pro Arg

<210> 947

<211> 36

<212> PRT

<213> Homo sapiens

<220>

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<222> -20..-1

<400> 947

Met Arg Lys Asp Val Arg Phe Leu Leu Phe Phe Thr Cys Gly Leu Pro -20 -15 -10 -5

Ala Leu His Gly Asp Ser Arg Val Glu Cys Ser Lys Ala His Pro Pro 1 5 10

Ala Met Tyr Tyr

<210> 948

<211> 48

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -27..-1

<400> 948

Met Leu Phe Trp Leu Pro Ser Pro Ser Glu Thr Thr Ser Ala Trp Thr
-25 -20 -15

Leu Leu Ser Ile Ser Leu Ser Val Phe Trp Ser Glu Pro Phe Asn Lys
-10 -5 1 5

Ser Leu Gly Ser Ser Lys Leu Pro Cys His Phe Phe Ser Ile Lys Arg
10 15 20

<210> 949

<211> 65

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

WO 99/53051 479 <222> -47..-1 <400> 949 Met Pro Val Cys Phe Tyr Ser Leu Ile Cys Phe Phe Ile Tyr Phe Cys -45 -40 -35 Leu Leu Ser Pro Arg Glu Thr Ile Glu Glu Val Ala Leu Phe Gln Phe -30 -25 -20 Ser Leu Leu Xaa Leu Gly Glu Gly Leu Thr Phe Leu Cys Leu Cys Gln -10 -5 Val Met Thr Asn Xaa Met Gln Leu Leu Phe Leu Ser Gly Val Val Cys 10 Gly <210> 950 <211> 21 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -13..-1 <400> 950 Met Ala Pro Leu Leu Ser Leu Ser Cys Ser Phe Ser Cys His Val -10 -5 Thr Leu Leu Pro Arg 5 <210> 951 <211> 47 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -20..-1 <400> 951 Met Val Pro Ala Ala Gly Ala Leu Leu Trp Val Leu Leu Leu Asn Leu -15 -10 Gly Pro Arg Ala Ala Gly Ala Gln Gly Leu Thr Gln Thr Pro Thr Glu Met Gln Arg Val Ser Leu Arg Phe Gly Gly Pro Met Thr Arg Arg 15 20 <210> 952 <211> 58 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -24..-1 <400> 952 Met Val Phe Trp Glu Ile Ser Val Gln Ile Ile Leu Ile Ser Glu Leu -20 -15

Leu Leu Leu Arg Ser Val Thr Ser His Asn Thr Met Met Arg Ala Leu -5 1 Ser Ser Gln Met Leu Ser Gln Ser Phe Pro Arg Pro Ser Phe Gly Phe 15 Ile Ser Lys Ile His Pro Ser His Pro Pro 30

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<211> 74
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -51..-1
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Met Phe Phe Leu Asn Ile Ala Met Phe Ile Val Val Met Val Gln Ile
                -45
Cys Gly Arg Asn Gly Lys Arg Ser Asn Arg Thr Leu Arg Glu Glu Val
                   ~30
                                     -25
Leu Arg Asn Leu Arg Ser Val Val Ser Leu Thr Phe Leu Leu Gly Met
              -15
                                  -10
                                              -5
Thr Trp Gly Phe Ala Phe Phe Ala Trp Gly Pro Leu Asn Ile Pro Phe
                      5
Met Tyr Leu Phe Ser Ile Phe Asn Ser Leu
  15
<210> 954
<211> 58
<212> PRT
<213> Homo sapiens
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<221> SIGNAL
<222> -17..-1
<400> 954
Met Asn Lys His Phe Leu Phe Leu Phe Leu Xaa Xaa Leu Ile Val
     -15
                          -10
                                   -5
Ala Val Thr Ser Leu Gln Cys Ile Thr Cys His Leu Arg Thr Arg Thr
                                      10
Asp Arg Cys Arg Arg Gly Phe Gly Xaa Cys Thr Ala Gln Lys Gly Glu
              20
Ala Cys Met Leu Leu Arg Ile His Gln Arg
<210> 955
<211> 47
<212> PRT
<213> Homo sapiens
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<221> SIGNAL
<222> -35..-1
<400> 955
Met Tyr Ile Lys Met Glu Ser Val Thr Leu Ser Pro Ala Pro Val Phe
                   -30
                                      -25
Pro Val Pro Ala Gln Leu Leu Leu Leu Thr Ser His Phe Leu Gly Glu
                                  -10
               -15
Ser Lou Gly Gly Gly Thr Leu Leu Val Pro Leu Leu Pro Pro Gly
                           5
<210> 956
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<213> Homo sapiens
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WO 99/53051 481 <220> <221> SIGNAL <222> -27..-1 <400> 956 Met Xaa Xaa Ala Leu Leu Arg Ser Arg Met Ile Gln Gly Arg Ile Leu -20 Leu Leu Thr Ile Cys Ala Ala Gly Ile Xaa Gly Thr Arg Gln Phe Gly -5 Tyr Asn Leu Ser Ile Ile Asn Asp 10 <210> 957 <211> 54 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -47..-1 <400> 957 Met Met Gly Xaa Leu Cys Pro Arg Ser Leu Pro Ile Pro Pro Met Ile -40 -45 Leu Ser Trp Trp Lys Met Gln Trp Lys Pro Leu Ala Leu Glu Asn Phe -25 Ser Gly Ser Cys Leu Phe Ser Xaa Ala Trp Leu Xaa Cys Xaa Cys His -10 Gly Asp Asp Asp Leu Ser 5 <210> 958 <211> 48 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -15..-1 <400> 958 Met Gly Leu Leu Gln Leu Leu Ala Phe Ser Phe Leu Gly Asn Ser Val -10 -5 1 Glu Thr Val Arg Gly Gly Gly Arg Thr Trp Ala Trp Gly Arg Lys Thr 10 Gln Lys Leu Leu Ala His Leu Arg Gly Ile Leu Gly Ala Trp Xaa Arg 25 <210> 959 <211> 25 <212> PRT <213> Homo sapiens <220> <221> SIGNAL

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<211> 48
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<213> Homo sapiens
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<221> SIGNAL
<222> -19..-1
<400> 960
Met Ser Phe Ser Ser Ala Leu Ile Leu Val Ile Ser Cys Leu Leu Leu
                -15
                                    -10
Ala Phe Glu Cys Val Cys Ser Cys Phe Ser Gly Ser Phe Asn Cys Asp
            1
                         5
                                                10
Val Arg Val Ser Ile Ser Asp Leu Ser Cys Phe Leu Leu Trp Gly Lys
                        20
   15
<210> 961
<211> 28
<212> PRT
<213> Homo sapiens
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<221> SIGNAL
<222> -22..-1
<400> 961
Met Gly Phe Trp Cys Gly Cys Pro Phe Cys Leu Xaa Val Phe Leu Leu
                            -15
Thr Asp Arg Thr Leu Ser Cys Arg Ser Val Gly Val
  -5
                        1
<210> 962
<211> 27
<212> PRT
<213> Homo sapiens
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<221> SIGNAL
<222> -15..-1
<400> 962
Met Val Leu Leu Ser Leu Ser Leu Trp Gly Ile Ser Thr Leu Ser Ser
                    -10
Thr Thr Ile Glu Leu Ile Tyr Thr Pro Ile Gly
<210> 963
<211> 28
<212> PRT
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<221> SIGNAL
<222> -25..-1
<400> 963
Met Ala Ser Leu Leu Ser Gly Phe Thr Ser Phe Cys Leu Leu His Val
                    -20
                                        -15
His Ser Phe Leu Pro Pro Val Phe Ser Thr Gln Asn
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-5

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<212> PRT
<213> Homo sapiens
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Met Glu Thr Ala Leu Xaa Xaa Thr Pro Gln Lys Arg Gln Val Met Phe
                   -25
                                        -20
Leu Ala Ile Leu Leu Xaa Xaa Trp Glu Ala Gly Ser Glu Ala Val Arg
               -10
                                    -5
Tyr Ser Ile Pro Glu Glu Thr Glu Ser Gly
                            10
<210> 965
<211> 66
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -35..-1
<400> 965
Met Met Leu Asp Phe Ala Leu Ser Pro Arg Leu Glu Arg Ser Gly Leu
                   -30
                                        -25
                                                            -20
Ile Met Ala Cys Cys Thr Leu Asp Leu Gly Ser Ser Pro Pro
                -15
                                    -10
Thr Ser Ala Ser Gln Val Ala Gly Thr Gly His Val Pro Pro His Pro
                           5
                                                10
Ala Ser Phe Phe Tyr Phe Xaa Val Xaa Gln Val Tyr Tyr Val Ser Gln
   15
                       20
Leu Ile
30
<210> 966
<211> 64
<212> PRT
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<222> -22..-1
<400> 966
Met Arg Thr Pro Gln Leu Ala Leu Leu Gln Val Phe Phe Leu Val Phe
                            -15
Pro Asp Gly Val Arg Pro Gln Pro Ser Ser Pro Ser Gly Ala Val
                       1
Pro Thr Ser Leu Glu Leu Gln Arg Gly Thr Asp Gly Gly Thr Leu Gln
               15
                                   20
Ser Pro Ser Glu Ala Thr Ala Thr Arg Pro Ala Val Pro Gly Leu Arg
           30
                                35
<210> 567
<211> 46
<212> PRT
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WO 99/53051 484 <222> -21..-1 <400> 967 Met Pro Arg Pro Arg Ala Cys Ala Ser Trp Pro Leu Leu Ala Ala Val -15 -10 Ser Gly Leu Arg Gly Leu Glu Trp Pro Pro Ser Trp Arg Arg Val Val Ala Ala Val Gly Val Cys Arg Val Arg Asp Trp Gly Pro Arg 20 <210> 968 <211> 23 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -17..-1 <400> 968 Met Asn Gly Ile Phe Leu Leu Leu Ile Ser Val Leu Thr Val Ile Trp -15 Phe Trp Lys Thr His Pro Gly 1 <210> 969 <211> 27 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -18..-1 <400> 969 Met Val Phe Leu Val Xaa Leu Leu Cys Ile Ile Xaa Leu Tyr Leu Ile -10 -15 -5 Arg Gly Ser Glu Trp Xaa Leu Pro Pro Asn Trp 5 <210> 970 <211> 53 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -18..-1 <400> 970 Met Met Thr Leu Ala Leu Phe Phe Leu Leu Arg Ile Ala Leu Ala Ser -15 -10 -5 Trp Ala Leu Phe Trp Ile His Met Asn Phe Arg Arg Ala Phe Phe His 5 10 Leu Arg Trp Phe Asp Ile Asn Ser Thr Glu Ser Val Asn Cys Phe Gly 20 25 Gln Tyi Gly Leu Ala

<210> 971 <211> 37 <212> PRT <213> Homo sapiens

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<222> -29..-1
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Met Ser Ile Arg Ser Asn Trp Ser Ser Val Glu Ser Lys Ser Arg Ile
                                    -20
Ser Leu Leu Val Phe Cys Leu Asn Asp Leu Ser Asn Ala Val Xaa Xaa
            -10
Gly Ile Glu Xaa Pro
   5
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<211> 120
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<400> 972
Met Ala Trp Ile Pro Leu Phe Leu Gly Val Leu Ala Tyr Cys Thr Gly
                        -10
                                            -5
Ser Val Ala Ser Tyr Glu Leu Thr His Pro Pro Ser Val Ser Val Ser
               5
                                    10
Pro Gly Gln Thr Ala Ser Ile Thr Cys Ser Gly Asp Lys Leu Gly Asp
           20
                                25
Lys Tyr Ala Cys Trp Tyr Gln Gln Lys Pro Gly Gln Ser Pro Val Leu
                            40
Val Ile Tyr Gln Asp Ser Lys Arg Pro Ser Gly Ile Pro Glu Arg Phe
                        55
                                            60
Ser Gly Ser Asn Ser Gly Asn Thr Ala Thr Leu Thr Ile Ser Gly Thr
                    70
                                        75
Gln Ala Met Asp Glu Ala Asp Tyr Tyr Cys Gln Ala Trp Asp Ser Ser
               85
                                    90
Thr Val Val Phe Gly Gly Gly Thr
           100
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<211> 32
<212> PRT
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<400> 973
Met Val Cys Val Ile Phe Lys Glu Leu Met Glu Phe Glu Phe Pro Gly
                -25 .
                            -20
Phe Cys Phe Xaa Leu Cys Phe Gly Arg Ser Ser Leu Cys Cys Arg Xaa
                                -5
<210> 974
<211> 78
<212> PRT
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<222> -30..-1
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Met Glu Ser Ser Gly Thr Pro Ser Val Thr Leu Ile Val Gly Ser Gly
                    -25
                                        -20
Leu Ser Cys Leu Ala Leu Xaa Thr Leu Ala Val Val Tyr Ala Ala Leu
                -10
                                    -5
Trp Arg Tyr Ile Arg Ser Glu Arg Ser Ile Ile Leu Ile Asn Phe Cys
                            10
Leu Ser Ile Ile Ser Ser Asn Ile Leu Ile Leu Val Gly Gln Thr Gln
                        25
Thr His Asn Lys Glu Tyr Leu His Asn His His Cys Ile Phe
                    40
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<211> 58
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<400> 975
Met Gly Val Cys Cys Ala Gln Asn Cys Ser Val Ser Gly Xaa Xaa Arq
   -30
                        -25
                                             -20
Asn Ala Leu Xaa Phe Leu Ala Ser Ser Phe Cys Phe Gly Glu Ala Asp
                    -10
                                        -5
Ser Gly Ser Arg Cys Cys Leu Lys Ile Ile Leu Gly Phe Tyr Leu Ile
                                10
Arg Tyr Ser Leu Ile Thr Tyr Gln Val Arg
       20
                            25
<210> 976
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<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -18..-1
<400> 976
Met Lys Ile Leu Tyr Leu Phe Phe Leu Lys Trp Ser His Pro Gly
            -15
                                -10
Trp Ser Ala Thr Xaa Trp Ser Trp His Thr Ala Thr Ser Ala Ser Leu
Ile Gln Val Ile Leu Pro Pro Trp
15
<210> 977
<211> 34
<212> PRT
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<221> SIGNAL
<222> -26..-1
<400> 977
Met Thr Pro Cys Phe Leu Gln Met Asp Asn Leu Thr Pro Leu Phe Leu
                        -20
                                            -15
Ser Gly Cys Phe Leu Phe Leu Ser Xaa Cys Xaa Ile Tyr Leu Ala Arg
-10
                                        1
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487
Ile Leu
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<211> 48
<212> PRT
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<222> -40..-1
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Met Gly Ser Ala Gly Arg Leu His Tyr Leu Xaa Met Thr Ala Glu Asn
                   -35
                            -30
Pro Thr Pro Gly Asp Leu Ala Pro Xaa Pro Leu Ile Thr Cys Lys Leu
                                   -15
                -20
                                                        -10
Cys Leu Cys Glu Gln Ser Xaa Gly Gln Asp Asp His Thr Pro Gly Met
                                1
<210> 979
<211> 88
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Met Asn His Leu Pro Pro Asn His Tyr Arg Xaa His Val Phe Thr Cys
                                   -40
His Val Asp Gln Tyr Leu Thr Val Glu Thr Ala Gly Gly Met Glu Lys
           -30
                               -25
                                                   -20
Glu Ala Val Ser Val Thr Val Leu Leu Ser Ala Ala Pro Cys Leu Leu
                           -10
Ser Cys Phe Leu Gly Ser Ser Val Ser Gly Leu Ala Phe Trp Val Ser
                                       10
Gln Gln Lys Thr Lys Gly Pro Glu Arg Cys Lys Asn Thr His His Xaa
               20
                                    25
Ala Xaa Asn Asn Phe Pro Ala Arg
<210> 980
<211> 42
<212> PRT
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<222> -40..-1
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Met Asn Lys Ile Lys Glu Asn Thr His Thr His Thr His Thr His Thr
                   ~35
                                       -30
His Lys Asn Asn Thr Lys Leu Val Ser Asn Leu Phe Leu Phe Met Leu
                                    -15
               -20
Pro Leu Trp Cys Ser Ile Gly Thr Cys Thr
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   -40
                   -35
                                  -30
Asn Leu Ile Tyr Leu Tyr Val Asp Ile His Ile His Lys Leu Phe Leu
                      -20
                                         -15
Tyr Ser Leu Phe Thr Glu Asn Val Leu Ala His Pro Cys Ile Val Leu
-10
Arg Arg Leu
<210> 982
<211> 37
<212> PRT
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<221> SIGNAL
<222> -33..-1
<400> 982
Met Gly Arg Leu His Arg Pro Arg Ser Ser Thr Ser Tyr Arg Asn Leu
                                          -20
        -30
                  -25
Pro His Leu Phe Leu Phe Leu Phe Val Gly Pro Phe Ser Cys Leu
                          -10
       -15
Gly Ser Tyr Ser Arg
  1
<210> 983
<211> 44
<212> PRT
<213> Homo sapiens
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<221> SIGNAL
<222> -27..-1
<400> 983
Met Gln Ser Gln Ala Ala Arg Glu His Lys Pro Gly Xaa Ser Arg Leu
                          -20
                                             -15
Leu Leu Leu Leu Leu Xaa Leu Pro Leu Pro Pro Pro Xaa Leu Arg
               -5
Thr Arg Xaa Phe Ser Xaa Thr Thr Leu Thr Ala Gly
              10
<210> 984
<211> 25
<212> PRT
<213> Homo sapiens
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<221> SIGNAL
<222> -15..-1
<400> 984
Met Arg Leu Trp Ser Leu Ala Cys Leu Ser Pro Pro Ala Val Gln Leu
                -10
                                     -5
Gly Ser Gln Gln Ala Thr Asp Trp Trp
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Met Ser Pro Leu Phe Ile Leu Ile Val Leu Ile Trp Ile Phe Ser Phe
-25
                    -20
                                        -15
Phe Phe Phe Ile Thr Leu Val Arg Gly Ser Ile Asn Leu Phe Phe
                                    1
<210> 986
<211> 25
<212> PRT
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<221> SIGNAL
<222> -22..-1
<400> 986
Met Asn Leu Gly Gly His Ser Asp His Ser Thr Phe Leu Phe Phe Leu
      -20
                            -15
Phe Phe Ser Val Phe Cys Phe Phe
                        1
<210> 987
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<222> -21..-1
<400> 987
Met Leu Asp Phe Ala Ile Phe Ala Val Thr Phe Leu Leu Ala Leu Val
                        -15
                                            -10
Gly Ala Val Leu Tyr Leu Tyr Pro Ala Ser Arg Gln Ala Ala Gly Ile
-5
                    1
Pro Gly Ile Thr Pro Thr Glu Glu Lys Asp Gly Asn Leu Pro Asp Ile
            15
                                20
Val Asn Ser Gly Ser Leu His Glu Xaa Leu Val Asn Leu His Glu Arg
                            35
Tyr Gly Pro Val Val Ser Phe Trp Phe Gly Arg Arg Leu Val Val Ser
                        50
Leu Gly Thr Val Asp Val Leu Lys Gln His Arg
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<400> 988

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<210> 992 <211> 89

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<222> -19..-1
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Met Glu Phe Gly Leu Ser Trp Val Phe Leu Val Ala Ile Ile Lys Gly
               -15
                                   -10
                                                       -5
Val Gln Cys Gln Val Gln Leu Val Glu Ser Gly Gly Leu Val Lys
Pro Gly Gly Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe
   15
Ser Asp Tyr Xaa Xaa Thr Xaa Ile Arg Xaa Ala Xaa Gly Lys Gly Leu .
                   35
                                       40
Xaa Trp Ile Xaa Xaa Ile Thr Thr Ser Gly Asn Thr Ala Xaa Tyr Ala
                                   55
Xaa Ser Val Lys Xaa Arg Phe Thr Ile
           65
<210> 993
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<222> -17..-1
<400> 993
Met Lys Arg Phe Phe Leu Phe Val Cys Leu Xaa Phe Asp Glu Ser Cys
       -15
                           -10
                                               -5
Ser Val Thr Arg Leu Gly Cys Cys Gly Ala Ile Ser Ala His Cys Xaa
                                      10
Leu Arg Leu Pro Gly Ser Ser Xaa Xaa Pro Ala Ser Thr Ser Arg Val
               20
                                   25
Xaa Gly Ile Thr Gly Met Arg
           35
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Pro Thr Lys His Ile Arg Glu His His Cys Met Leu Phe Val Ser Phe
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Leu Leu Leu Leu Gly Ser Arg
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<221> SIGNAL

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Met Leu Ser Cys Cys Gln Gly Ala Cys Cys Pro Ser Thr Pro His Glu
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Gly Thr Arg Ser Thr Val Ser Trp Ile Pro Pro Thr Tyr Lys Ala Ala
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Thr Gln
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Met Val Arg Ala Ser Ile Leu Leu Ser Met Phe Cys Val Ser His Thr
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Val Gln Thr Ala Thr Tyr Thr
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       -15
Cys Ala Pro Arg Thr Tyr Val Val Ser Ala Thr Thr Leu Ser Ala Val
                                       10
Gln Gly His Cys Pro Leu Gln Ser Arg Thr Ser Thr Lys Gly Lys Leu
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Trp Pro Phe Gly
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<400> 998
Met Ile Phe Thr Phe Gln Gln Ile Gly Gly Lys Leu Leu Ser Gly
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           -20
                                                   -10
Leu Thr Gln Glu Cys Leu Gly Ala Leu Pro Glu Ala Asn Val Phe Cys
     ~5
                           1
Arg Gly Gly Cys Thr Ala Thr Val Leu Lys His Gly Lys Ala Ser Pro
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Glu Ser
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<211> 46
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<400> 999
Met Asn Cys Val Arg Gln Ala Asn Ile Arg Met Gln Cys Lys Ile Tyr
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Asp Ser Leu Leu Ala Leu Ser Pro Asp Leu Gln Ala Ala Arg Gly Leu
                   -10
                                -5
Met Cys Ala Ala Ser Val Met Ser Phe Leu Ala Phe Met Met
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                           -30
Asp Phe Cys Thr Leu Thr Leu Tyr Pro Gly Thr Leu Leu Lys Leu Leu
               -20
                                   -15
Ile Ser Leu Arg Ser Phe Trp Ala Glu Thr Thr Gly
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<211> 43
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<213> Homo sapiens
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Met Phe Ser Ser Pro Gly Leu Arg Thr Leu Phe Val Leu Val Gly Ser
-25
                   -20
                                     -15
Leu His Leu Phe Leu Ser Val Leu Ala Ser Lys Ser Arg Asn Ser Lys
               - 5
                                                 5
                                  1
Lys Gln Arg Leu Phe Leu Leu Val Pro Leu Tyr
       10
                           15
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Ala Pro Gly Leu Cys Ser Gly Gln Pro Gly Val Arg Ala Trp Pro Gly
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                            1
Val Thr Gln Leu Thr Gln Xaa Glu Glu Cys Pro Trp Phe Ser Ala Leu
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Glu Gly Leu
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                                                    -20
Gly Pro Leu Leu Leu Leu Ser Leu Ile Phe Gly Pro Cys Ile
                            -10
                                                -5
Leu Asn Ser Phe Leu Asn Xaa Ile Lys Gln Arg Ile Ala Ser Gly Lys
Arg
<210> 1004
<211> 102
<212> PRT
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<400> 1004
Met Ala Gly Ser Arg Gln Arg Gly Leu Arg Ala Arg Val Arg Pro Leu
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                                    -20
Phe Cys Ala Leu Leu Ser Leu Xaa Xaa Xaa Pro Xaa Xaa Arg
           -10
                                -5
Arg Xaa Arg Arg Pro Arg Gly Arg Val Ala Thr Ser Pro Phe Arg Val
                        10
Xaa Ile Gln Leu Gln Gly Ala Ala Pro Gly Ala Glu Arg Arg Asp Arg
                    25
                                        30
Ala Leu Leu Gly Pro Arg Gly Glu Cys Tyr Ser Lys Phe Arg Ser Asn
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Ser Ser Ser Thr Ile Phe Lys Lys Xaa Lys Arg Leu Ser Val Xaa Xaa
Asp Xaa Ser Gly Pro Gly
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Val His Cys Asp Val Gln Leu Val Glu Ser Gly Gly Leu Val Gln

495 10 Pro Gly Gly Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Leu Thr Leu 20 25 Ser Asn Asp Trp Met His Trp Val Arg Gln Ala Pro Gly Lys Gly Leu 40 Val Trp Val Ser His Ile Asp Ser Ser Xaa Thr Ile Thr Asn Tyr Ala 55 Asp Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Trp 70 <210> 1006 <211> 38 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -15..-1 <400> 1006 Met Gly Leu Phe Leu Gly Phe Leu Ala Cys Ser Val Ala Tyr Gln Cys -10 -5 His Ser Ala Phe Val Thr Val Ala Ser Gln Tyr Thr Leu Lys Ser Glu Thr Leu Met Pro Ala Ala 20 <210> 1007 <211> 104 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -49..-1 <400> 1007 Met Trp Glu Asp Ser Arg Asn Lys Arg Gly Gly Arg Trp Leu Val Ser -45 -40 Leu Ala Lys Gln Gln Arg His Ile Glu Leu Asp Arg Leu Trp Leu Glu -30 -25 -20 Thr Phe Ser Val Phe Leu Gly Leu Ile Phe Phe Leu Glu Leu Ala Thr -10 -5 Gly Ile Leu Ala Phe Val Phe Lys Asp Trp Ile Arg Asp Gln Leu Asn 10 Leu Phe Ile Asn Asn Asn Val Lys Ala Tyr Arg Asp Asp Ile Asp Leu 25 Gln Xaa Leu Ile Asp Phe Ala Gln Glu Tyr Trp Ser Cys Cys Gly Xaa 35 Glu Ala Pro Ile Xaa Gly Thr Gly 50 <210> 1008 <211> 34 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -14..-1

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60

Thr Leu Asn Phe Gly Pro Gln His Pro Ala Ala His Gly Val Leu Arg

55

75 80 70 Leu Val Met Glu Leu Ser Gly Glu Met Val Arg Lys Cys Asp Pro His 90 95

Ile Gly Leu Leu His Arg Gly Thr Glu Lys Leu Ile Glu Tyr Lys Xaa 105 110

Tyr Leu Gln Ala Leu Pro Tyr Phe 115 120

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<220>

<221> SIGNAL

<222> -28..-1

<400> 1012

Met Leu Ile Trp Ser Ser Ser Phe Pro Ala Pro Pro Leu Phe Leu -25 -20 -15

Val Phe Leu His Leu Phe Leu Xaa Val Tyr Leu Gly Leu Val Met Pro -10 -5

Thr Gln Gln Tyr Leu Leu Gln Ser Pro Leu Met Phe Thr Asp Lys 15

Ala Gln

<210> 1013

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Met Cys Arg Met Cys Arg Phe Val Thr Trp Ile Asn Val Cys His Gly -40 -35

Asp Leu Leu His Arg Ser Ser Arg Arg Leu Gly Val Lys Pro Ser Thr -30 -25 -20

His Trp Leu Phe Phe Leu Met Leu Ser Leu Cys Thr Pro Pro Asp Arg -10

Pro Trp Cys Val Leu Phe Pro Pro Leu

<210> 1014

<211> 40

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -31..-1

<400> 1014

Met Xaa Thr Gln Glu Ala Gly Leu Ile Phe Phe Ser Pro Pro Phe Ser

-25 -20

Leu Ser Leu Ser Leu Pro Leu Ser Leu Xaa Leu Leu Xaa Xaa -10 -5

Pro His Ser Arg Thr Pro Gln Arg

<221> SIGNAL

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Gln Ala Trp Arg Leu Met Pro Val Val Pro Ala Val Trp Glu Pro Glu
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Ala Gly Gly Leu Leu Gln Leu Gly Gly Ser Arg
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<211> 88
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                               -25
   -35
                          -30
Ala Val Asp Ala Ile Pro Phe Cys Leu Leu Val Phe Phe Leu Ile Val
                      -15
                                          -10
Arg Thr Leu Ser Cys Arg Ser Val Gly Val Cys Trp Arg Ser Thr Pro
                   1
                                  5
Asp Pro Val Cys Leu Gly Ile Thr Ser Arg Gly Cys Arg Thr Glu Ile
           15
                              20
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Ser Gln Arg Gly Thr Glu Cys Met
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                           -10
Ala Asn Ala Gln Ser Lys Phe Ser Leu Tyr Phe Phe Pro Leu Val Lys
Pro Gly
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<400> 1031

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Are Mot The Luc Ace Dro Lou Clu

Met Arg Met Thr Lys Asp Pro Leu Gly Ser Leu Ile Ala Ser Leu Ala
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-15
-5
Pro Ser Thr Gly Leu Gly

<210> 1032

<211> 57

<212> PRT

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<220>

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<222> -28..-1

<400> 1032

Met Lys Leu Gln Phe Ala Phe Cys Tyr Phe Leu Tyr Leu Asp Thr Phe
-25 -20 -15

Phe Leu Phe Leu Phe Phe Xaa Glu Xaa Xaa Xaa Xaa Xaa Xaa Gly
-10 -5 1

Arg Ser Ala Val Ala Xaa Pro Gln Leu Xaa Ala Ala Ser Thr Phe Xaa 5 10 15 20

Phe Gln Ala Ile Phe Leu Pro Gln Xaa

25

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<222> -69..-1

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Met Ala Ala Gly Glu Leu Glu Gly Gly Lys Pro Leu Ser Gly Leu Leu
-65 -60 -55

Asn Ala Leu Ala Gln Asp Thr Phe His Gly Tyr Pro Gly Ile Thr Glu
-50 -45 -40

Glu Leu Leu Arg Ser Gln Leu Tyr Pro Glu Val Pro Pro Glu Glu Phe
-35 -30 -25

His Pro Phe Leu Ala Lys Met Arg Gly Ile Leu Lys Val Leu Leu Phe
-20 -15 -10

Ser Val Val Ser Gly Leu Glu Gln Asn Pro Leu Ala Ala Gly Phe Arg -5 5 10

Leu Ser His Pro

15

<210> 1034

<211> 47

<212> PRT

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<221> SIGNAL

<222> -31..-1

<400> 1034

Met Met Met Ser Asn Val Met Leu Met Leu Gln Leu Gln Pro Leu Leu
-30 -25 -20

Ala Xaa Ser Leu Ile Leu Ser Pro Ser Pro Arg Pro Val Leu Gly Phe
-15 -5 1

Phe Arg Gln Val His Leu Leu Thr Arg Ser His Phe Ser Arg Trp
5 10 15

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. <222> -20..-1
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               -15
                                       -10
 Pro Ile Ser Ala Ser Ser Asn Tyr His Phe Thr Leu Tyr Leu His Asp
               1
                               5
 Ile Asn Phe Phe Ser
     15
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             -10
                         -5
 -15
 Thr Arg
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 <211> 25
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 <222> -13..-1
 <400> 1037
 Met Gly Leu Phe Leu Cys Cys Ser Leu Leu Ile Phe Cys Leu Val Val
                    -5
            -10
 Leu Ile Ile Thr Glu Leu Gly Tyr Gly
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 <211> 30
 <212> PRT
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 <222> -14..-1
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                -10
 Arg Val Pro Phe Cys Ser Trp Glu Lys Ser Asp Gly Arg Ser
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<210> 1039

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                           -35
     -40
Asp Gly Phe Leu Lys Tyr Ser Asp Pro Asn Asp Ile Ala Leu Ser Val
                       -20
Leu Ser Leu Val Ile Asn Phe Ser Trp Ser Arg Lys Cys Phe Val Pro
                <del>-</del>5
Tyr Tyr Ile Pro Phe Lys Pro Tyr Arg Xaa Pro Tyr Pro Thr Ala Ala
                                15
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Arg
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Met Tyr Val Cys Ile Tyr Ile Xaa Leu Xaa Asp Leu Tyr Asp Phe Phe
                -35
                                  -30
Leu Leu Gly Thr Tyr Phe Phe Glu Arg Lys Cys Phe Val Cys Xaa Leu
                               -15
                                                    -10
Phe Val Phe Leu Leu Ser Gly Leu Asn Tyr Phe Ser Ile Leu Ser Phe
        -5
Tyr Pro Arg
<210> 1041
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<222> -40..-1
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Met Cys Ile Phe Cys Leu Phe His Leu Leu Tyr His Lys Leu Leu Ser
                                         -30
               . -35
Arg Ser Leu Phe Phe Cys Cys Ile Phe Ser Gly Phe Ile Thr Phe Ile
                                     -15
                -20
 Phe Ser Phe Ser Phe Cys Glu Cys Ile Val Gly Met Tyr Ile Tyr Gly
 Ala Arg
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<400> 1042
Met Xaa Ile Cys Tyr Asn Ile Phe Gln Asn Ile Leu Gly Leu Leu
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     -25
Ile Phe Leu Tyr Leu Ser Leu Asn Leu Phe Cys Ile Phe Phe Ser Val
                                         1
                      -5
Pro Ala Leu Gln Pro Arg Arg Leu
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<210> 1043
<211> 29
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<222> -26..-1
<400> 1043
Met Ala Ser Ser Met Leu Xaa Ser Phe Gln Thr Phe Met Met Leu Thr
                                           -15
                       -20
Leu Leu Gly Phe Pro Ser Lys Ala Leu Thr Phe Ile Ser
                   -5
<210> 1044
<211> 33
<212> PRT
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<400> 1044
Met Gly Arg Ser Lys Arg Gln Leu Leu Ser Leu Pro Gly Ser Phe Ile
                                       -10
                -15
Pro Gly Asn Cys Arg Pro Arg Ile Leu Ser Asn Gly Glu Xaa Arg Arg
                1
Lys
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 <222> -25..-1
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 Met Arg Ser Asp Gly Phe Ile Arg Gly Phe Cys Phe Cys Phe Phe Leu
                                        -15
                    -20
 Ile Phe Leu Leu Pro Pro Leu Pro Ala Met Ile Leu Arg Pro Leu Gln
                                   1
                - 5
 Pro Cys Gly Ile Ile Ser Pro Ile Lys Pro Leu Phe Pro Phe Phe
                                               20
                            15
        10
 <210> 1046
 <211> 39 ·
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<213> Homo sapiens

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                   -10
                                   -5
Gln Arg Thr Met Ile His Trp Asn Val Phe Leu Trp Asn Ser Phe Tyr
              5
                                 10
Ser Cys Ile Lys Ile Phe Pro
           20
<210> 1047
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<213> Homo sapiens
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<221> SIGNAL
<222> -31..-1
<400> 1047
Met Thr Trp Thr Lys Cys Pro Leu Pro Leu Gly Pro Ala Phe Phe Thr
                      -25
Gln Cys Cys Leu Ile Gly Leu Leu Val Pro Leu Leu Gly Trp Gly Asn
        -10
Gln Asn Thr Gln Trp Tyr Pro Thr Ser Lys Met Pro Asp Gly
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Gly Trp Glu Ile Val Tyr Phe Leu Val Leu Leu Arg Val Leu Tyr Thr
   -15
                      -10
Leu Gln Trp Gly Gly
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Ser Ser Ser Cys Ser Ile Ala Pro
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Met Arg Phe Arg Phe Cys Gly Asp Leu Asp Cys Pro Asp Trp Val Leu
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                -75
Ala Glu Ile Ser Thr Leu Ala Lys Met Ser Ser Val Lys Leu Arg Leu
                                -55
            -60
Leu Cys Ser Gln Val Leu Lys Glu Leu Leu Gly Gln Gly Ile Asp Tyr
                                                -35
                           -40
Glu Lys Ile Leu Lys Leu Thr Ala Asp Ala Lys Phe Glu Ser Gly Asp
                                            -20
                        -25
Val Lys Ala Thr Val Ala Val Leu Ser Phe Ile Leu Ser Ser Ala Ala
                                        -5
                   -10
Lys His Ser Val Asp Gly Glu Ser Leu Ser Ser Glu Leu Gln Gln Leu
                               10
                                                    15
Gly Leu Pro Lys Glu His Ala Ala Ser Leu Cys Arg Cys Tyr Glu Glu
                                                30
                           25
Lys Gln Ser Pro Leu Gln Lys His Leu Arg Val Cys Ser Leu Arg Met
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Asn Arg
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Met Phe Leu Ala Ala Leu Phe Thr Val Ala Lys Ile Trp Lys Gln Pro
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                -10
Lys Cys Ser Ser Thr Asn Lys Trp Thr Lys Lys Met Trp Tyr Ile Tyr
                            10
 Thr Met Glu Tyr Tyr Ser Ala Ile Lys Lys Asp Asp Ile Leu Ser Phe
                        25
 Ala Thr Ile Trp Met Glu Leu Glu Ser Ile Thr Leu Ser Glu Ile Ser
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 Gly Xaa Pro Lys Asp Lys Leu Leu Met Phe Ser Leu Ile Cys Gly
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<222> -27..-1

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<210> 1053

509

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                              -70
Pro Gly Pro Phe Thr Gln Tyr Leu Val Asp His His His Thr Leu Leu
  -60
                          -55
                                              -50
Cys Asn Gly Tyr Trp Leu Ala Trp Leu Ile His Val Gly Glu Ser Leu
                  -40
                                          -35
Tyr Ala Ile Val Leu Cys Lys His Lys Gly Ile Thr Ser Gly Arg Ala
                  -25
                                      -20
Gln Leu Leu Trp Phe Leu Gln Thr Phe Phe Phe Gly Ile Ala Ser Leu
              -10
Xaa Ile Leu Ile
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Met Cys Cys Trp Ile Trp Val Ala Ser Ile Leu Leu Arg Ile Phe Ala
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           -10
Ser Val Leu Ile Arg Asp Ile Tyr Leu Trp Phe Ser Phe Phe Phe
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Met Ile Ser Ser His Leu Tyr Asn Phe Ser Leu Leu Phe Phe Xaa Leu
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           -20
Trp Leu Arg Tyr Lys Glu Ser Gly Arg Glu Gly Asn Cys Glu Glu Gly
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Ala Phe Ser Arg Trp
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Gly Ala Ser Asn Ala Thr Arg Xaa Pro Lys Xaa Leu Tyr Arg Xaa Tyr
                                       10
Asn His Gly Val Leu Lys Ile Thr Ile Cys Lys Ser Cys Gln Lys Pro
               20
                                   25
Val Asp Lys Tyr Ile Glu Tyr Asp Pro Val Ile Ile Leu Xaa Asn Ala
                               40
Ile Leu Cys Lys Ala Xaa Ala Tyr Arg His Ile Leu Phe Asn Thr Gln
                           55
Ile Asn Asn Lys Leu Pro Ile Leu Leu Ala Phe Leu Pro Ser Cys Gly
Xaa Thr Ala His Asp Gly Lys Lys Pro Asn Phe Ile Leu Leu Leu
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Lys Xaa Tyr Tyr Tyr Leu Ala Thr Glu Asn
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Met Ala Ala Gly Val Ser Leu Leu Ala Leu Val Val Arg Val Île Leu
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Ser Thr Ala Ile Leu Cys Pro Ser Gly Ala Ser Arg Arg Gln Arg Ser
Ser Glu Val Glu Trp Gly Thr Asp Ser
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Met Asn Pro Leu Phe Trp Leu Ile Leu Cys Ser Gly Leu Leu Cys Asn
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-15
                 -10
Lys Ser Phe
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-10

-15

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Leu Ser Leu Phe
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          -20
Pro His Phe Phe Leu Ser Phe Leu Ser Pro Phe Tyr Leu His Pro Trp ,
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Leu Phe Lys Phe Leu Ala His Phe Leu Ile Gly Leu Thr Val Cys Phe
                         -10
Gly Glu Gly Xaa Leu Met Ser Tyr Arg Ser Ser Tyr Leu Leu Lys
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Gly Pro Pro Gly
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Met Gly Phe Trp Cys Glu Cys Pro Phe Cys Leu Leu Val Phe Leu Leu
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  -20
Thr Glu Trp Thr Ser Ser Lys Leu Gln Lys Thr
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 Met Trp Trp Gly Arg Cys Phe Ile Arg Val Leu His Leu Phe Pro Leu
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-15

-20

-10

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Thr Pro Ala Ser Thr Gly His Trp
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Met Arg Asp Pro Leu Ala Asp Met Val His Ser Tyr Leu Ser Ser Ser
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             . -25
Leu Phe Met Ala Leu Pro Pro Val Leu Ser Ser His Gly Ser Arg Asn
            -10
                                ~5
Leu Arg Ile Trp Gly Ser Pro Phe Gly Gly Ala Leu Thr Lys Gly Lys
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Ala Pro Pro Thr Pro Ala Gln Pro Ala Leu
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Met Ser Ser Ala Trp Leu Cys Leu Pro Cys Ser Leu Cys Val Ser Gln
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    -15
Leu Leu Pro Ser Tyr Ser Leu Leu Ile Pro Ala Pro
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 Pro Pro Met Arg Ala Cys Ser Val Cys Val Leu
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513 -10 -15 Phe Ala Asn Pro Lys Leu Ser Gly Pro Ile Ser Ile Ser Val Thr Ser Ala Gly Ser Pro Pro Gly Ala <210> 1068 <211> 26 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -15..-1 <400> 1068 Met Lys Asp Leu Leu Gly Thr Ala Phe Leu Glu Gly Ser Leu Ala Ala Tyr Leu Thr Met Ala Asn Ile Thr His Val <210> 1069 <211> 29 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -19..-1 <400> 1069 Met Ala Asn Asp Ile Lys His Leu Phe Met Cys Leu Leu Thr Ile Cys -15 Ile Ser Ser Leu Glu Lys Leu Pro Phe Phe Phe Phe <210> 1070 <211> 98 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -24..-1 <400> 1070 Met Tyr Gln Lys Val Thr Ser Tyr Cys Arg Ser Ala Thr Leu Val Gly -20 -15 Phe Thr Val Gly Ser Val Leu Gly Gln Ile Leu Val Ser Val Ala Gly -5 Trp Ser Leu Phe Ser Leu Asn Val Ile Ser Leu Thr Cys Val Ser Val 15 Ala Phe Ala Val Ala Trp Phe Leu Pro Met Pro Gln Lys Ser Leu Phe 30 Phe His His Ile Pro Ser Thr Cys Gln Arg Val Asn Gly Ile Lys Val 50 4.5 Gln Asn Gly Gly Ile Val Thr Asp Thr Gln Leu Leu Thr Pro Ser Trp 65 Leu Gly <210> 1071 <211> 19 <212> PRT

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Gly Leu Phe
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Met Asn Cys Val Thr Leu Ile Gln Ala Leu Ser Leu Trp Ala Ser Val
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                       -15
                                           -10
Ser Pro Ser Trp Met Cys Arg Pro Pro Ala Ser Phe Ile Ile Thr Thr
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Thr Thr Thr Cys Gly
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 -15
Ser Pro Arg
<210> 1074
<211> 255
<212> PRT
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                                -25
Val Thr Ala Giu Lys Met Ala Val Leu Ala Pro Leu Ile Ala Leu Val
                               -10
           -15
Tyr Ser Val Pro Arg Leu Ser Arg Trp Leu Ala Gln Pro Tyr Tyr Leu
                                          10
Leu Ser Ala Leu Leu Ser Ala Ala Phe Leu Leu Val Arg Lys Leu Pro
                   20
                                       25
Pro Leu Cys His Gly Leu Pro Thr Gln Arg Glu Asp Gly Asn Pro Cys
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35 40 45 Asp Phe Asp Trp Arg Glu Val Glu Ile Leu Met Phe Leu Ser Ala Ile 55 Val Met Met Lys Asn Arg Arg Ser Ile Thr Val Glu Gln His Ile Gly 70 Asn Ile Phe Met Phe Ser Lys Val Ala Asn Thr Ile Leu Phe Phe Arg 85 Leu Asp Ile Arg Met Gly Leu Leu Tyr Ile Thr Leu Cys Ile Val Phe 100 105 Leu Met Thr Cys Lys Pro Pro Leu Tyr Met Gly Pro Glu Tyr Ile Xaa 115 120 Tyr Phe Asn Asp Lys Thr Ile Asp Glu Glu Leu Glu Arg Asp Lys Arg 130 135 Val Thr Trp Ile Val Glu Phe Phe Ala Xaa Trp Ser Asn Asp Cys Gln 150 Ser Phe Ala Pro Ile Tyr Ala Asp Leu Ser Leu Lys Tyr Asn Cys Thr 165 Gly Leu Asn Phe Gly Lys Val Asp Val Gly Arg Tyr Thr Asp Val Ser 180 185 Thr Arg Tyr Lys Val Ser Thr Ser Pro Leu Thr Lys Gln Leu Pro Thr 195 200 Leu Ile Leu Phe Gln Gly Gly Lys Glu Ala Met Arg Arg Pro Gln 215

<210> 1075

<211> 153

<212> PRT

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<220>

<221> SIGNAL

<222> -17..-1

<400> 1075

 Met
 Thr
 Met
 Tyr
 Leu
 Trp
 Leu
 Lys
 Leu
 Ala
 Phe
 Gly
 Phe
 Ala
 Phe
 Phe</th

80 85 90 95.
Ala Asp Ser Gln Thr Pro Ser Ala Gly Thr Asp Thr Gln Thr Phe Ser
100 105 110

Gly Ser Ala Xaa Met Gln Asn Ser Thr Leu Pro Gln Ala Ala Met Leu 115 120 125

Ser Gln Met Ser Gln Glu Arg Gly Val 130 135

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<211> 42

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<222> -17..-1

WO 99/53051 516 <400> 1076 Met Thr Met Tyr Leu Trp Leu Lys Leu Leu Ala Phe Gly Phe Ala Phe -15 -10 Leu Asp Thr Glu Val Phe Val Thr Gly Gln Ser Pro Thr Pro Ser Pro 5 Thr Gly Val Ser Ser Val Gln Thr Pro Gln 20 <210> 1077 <211> 87 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -17..-1 <400> 1077 Met Thr Met Tyr Leu Trp Leu Lys Leu Leu Ala Phe Gly Phe Ala Phe -15 -10 Leu Asp Thr Glu Val Phe Val Thr Gly Gln Ser Pro Thr Pro Ser Pro 10 Thr Gly Val Ser Ser Val Gln Thr Pro His Leu Pro Thr His Ala Asp 20 Ser Gln Thr Pro Ser Ala Gly Thr Asp Thr Gln Thr Phe Ser Gly Ser 40 Ala Xaa Met Gln Asn Ser Thr Leu Pro Gln Ala Ala Met Leu Ser Gln 50 Met Ser Gln Glu Arg Gly Val 65 <210> 1078 <211> 42 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -36..-1 <400> 1078 Met Arg Gly Ala Thr Trp Pro Trp Pro Cys Leu Pro Ala Arg Thr Ser -30 -25 Thr Ala Ala Ser Ile Ala Arg Leu Phe Leu Leu Ser Gly Thr Ile Trp -15 Ile Ala Ile Cys Lys Pro Thr Thr Asn Gly <210> 1079 <211> 72 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -64..-1

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Ile Asn Arg His Leu Ile Tyr Cys Leu Gly Asp Ile Ile Leu Xaa Xaa

-30 -25 -20

Leu Asp Leu Ser Ala Leu Leu Arg Ser Leu Leu Pro Xaa Leu Xaa
-15 -5

Gln Ile Pro Gln Ala Thr Leu Arg

<210> 1080

<211> 42

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -15..-1

<400> 1080

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Leu Thr Val Met Val Glu Gly Lys Glu Glu Gln Val Thr Ser Tyr Thr 5 10 15

Asp Gly Ser Arg Gln Arg Asp Ser Asn Phe 20 25

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<211> 64

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<220>

<221> SIGNAL

<222> -39..-1

<400> 1081

Met Lys Arg Ile Arg Arg Lys Arg Arg Asn Glu Val Thr Ile Gln Pro
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Phe Pro Ile Arg Leu Pro Leu Leu Pro Pro Leu Ile Ser Phe Leu His
-20 -15 -10

Thr Leu Gln Val Val Cys Ser Val Ile Met Lys Ser Ile Arg Lys Ala

Phe Val Leu Cys Gly Phe Leu Tyr Phe Glu Phe Phe Asp Gln Lys Leu 10 20 25

<210> 1082

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<212> PRT

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<222> -22..-1

<400> 1082

Met Leu Pro Leu Leu His Cys Phe Phe Xaa Val Xaa Leu Phe Xaa Xaa -20 -15 -10

Val Xac Val Xaa Xaa Ala Ala Leu Leu Arg Tyr Asn Xaa Ser Ile Gln
-5 1 5 10

Xaa Gly Arg Ala Gln Xaa Leu Xaa Pro Xaa Ile Pro Xaa Leu Trp Glu 15 20 25

Thr Lys Xaa Gly Arg Leu Leu Glu Pro Arg Asn

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518 <211> 30 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -21..-1 <400> 1083 Met Val Ser Val Phe Arg Ser Glu Glu Met Cys Leu Ser Gln Leu Phe -20 -15 -10 Leu Gln Val Glu Ala Ala Tyr Cys Cys Val Ala Glu Leu Gly 1 5 <210> 1084 <211> 41 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -28..-1 <400> 1084 Met Ala Ala Leu Arg Ser Thr Leu Thr Trp Thr Glu Val Val Gly Trp -25 -20 -15 Trp Ser Val Ala Ser Leu Leu Ser Asp Val Ala Ala Trp Trp Pro Pro -10 -5 1 His Ser Thr Ser Thr Arg Gly Gly Val <210> 1085 <211> 47 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -44..-1 <400> 1085 Met Asn Ala Leu Val Asp Gly Lys Arg Leu Xaa Xaa Cys Ile Arg Tyr -40 -35 -30 Phe Asp Ser Ile Ser Leu Tyr Ser Lys Ala Ser Leu Ser Cys Cys Leu -25 -20 -15 Val Cys Val Phe Thr Cys Ser Leu Leu Ala Phe Phe Ser Pro Cys -5 <210> 1086 <211> 84 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -19..-1 <400> 1086 Met Glu Phe Gly Leu Ser Trp Val Phe Leu Val Ala Ile Leu Lys Gly -15 -10 Val Gln Cys Glu Leu Gln Val Val Glu Ser Gly Gly Leu Val Gln

Pro Gly Arg Ser Leu Arg Leu Ser Cys Arg Thr Ser Gly Phe Ala Phe

519 20 25 Asp Asp Tyr Asn Leu Ser Trp Val Arg Gln Ala Pro Gly Lys Gly Leu 35 40 45 Glu Trp Val Gly Phe Ile Arg Ser Lys Pro Tyr Gly Glu Thr Thr 55 Tyr Ala Ala Trp <210> 1087 <211> 19 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -14..-1 <400> 1087 Met Ser Leu Phe Xaa Leu Xaa Xaa Leu Arg Gln Ser Phe Thr Xaa Xaa -10 Ala Gln Ala 5 <210> 1088 <211> 30 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -19..-1 <400> 1088 Met Ile Ser Ala His Cys Ser Phe Tyr Phe Leu Ala Ser Ser Ser Leu -15 -10 Ser Thr Ser Ala Ser Xaa Arg Thr Gly Ile Thr Asp Val Ser 5 <210> 1089 <211> 43 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -24..-1 <400> 1089 Met Asn Ala Glu Asn Asn Phe Phe Gly Phe Val Cys Leu Phe Val Phe -20 -15 Leu Tyr Thr Thr Pro Cys Asn Cys Phe Gly Leu Glu His Leu Trp Ile Leu Ser Phe Met Val Val Leu Gly Xaa Thr Arg <210> 1090 <211> 31 <212> PRT <213> Homo sapiens <220> <221> SIGNAL

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                                -15
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Leu Leu Thr Cys Val Ser Ser Leu Arg Xaa Asp Ile Tyr Pro His
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<211> 34
<212> PRT
<213> Homo sapiens
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<221> SIGNAL
<222> -25..-1
<400> 1091
Met Arg Arg Lys Arg Arg Glu Arg Lys Glu Arg Lys Ser Ile Leu Leu
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                                     -15
Ala Ala Leu Ser Arg Asn Ile Ser Pro Gly Gln Thr Tyr Arg Thr Ser
               -5
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Pro Ala
<210> 1092
<211> 30
<212> PRT
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<221> SIGNAL
<222> -23..-1
<400> 1092
Met Gly Ser Pro Tyr Val Ala His Val Gly Leu Glu Leu Leu Thr Ser
        -20
                               -15
Ser Asp Pro Pro Ser Leu Ala Ser Gln Val Leu Gly Ile His
                            1
<210> 1093
<211> 45
<212> PRT
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<222> -19..-1
<400> 1093
Met His Leu Tyr Thr His Val Cys Trp Leu Thr Leu Thr Leu Ala His
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Ser His Ser Leu Thr His Thr His Thr Leu Thr Pro Ser His Thr Arg
Thr His Ser His Thr Cys Ala Cys Leu His Ala His Lys
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:210> 1094
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<222> -14..-1
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Pro Ser Gly
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<210> 1098
<211> 38
<212> PRT
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<222> -21..-1
<400> 1098
Met Gly Ser Arg Pro Val Ser Xaa Ala Gly Leu Glu Leu Leu Ala Ser
                        -15
                                             -10
Ser Asn Ser Ser Ala Leu Pro Phe Gln Cys Ser Gly Ile Thr Gly Met
                    1
Ser Xaa His Thr Leu Ala
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<210> 1099
<211> 19
<212> PRT
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<221> SIGNAL
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<400> 1099
Met Leu Cys His Leu Ser Leu Val Phe Leu Gly Xaa Gly Gln Phe Trp
Ser Gln Asn
   5
<210> 1100
<211> 30
<212> PRT
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<220>
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<222> -17..-1
Met Thr Asn Leu Phe Met Cys Leu Phe Ala Ile Cys Ile Ser Ser Asn
                            -10
Ala Lys Cys Leu Phe Ser Leu Phe Pro Phe Phe 11e Giu Gly
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<210> 1101
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                                                -15
Leu Phe Thr Leu Leu Val Ser Thr Arg Ser Gly Arg Ser Arg Ala Gly
                     -5
Cys Ala Trp Arg Trp Arg Gly Arg Trp Ser Val Gly Gln Lys Gly Xaa
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                                    15
<210> 1102
<211> 28
<212> PRT
<213> Homo sapiens
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<221> SIGNAL
<222> -15..-1
<400> 1102
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Gln Ala Thr Arg Ala Thr Thr Pro Cys Arg Leu Arg
<210> 1103
<211> 41
<212> PRT
<213> Homo sapiens
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<221> SIGNAL
<222> -37..-1
<400> 1103
Met Cys His Arg Arg Trp Leu His Leu Ser Thr Arg His Leu Gly Phe
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Lys Pro Arg Ile His Tyr Val Phe Val Leu Met Leu Ser Leu Pro Leu
                       -15
Pro Pro Thr Pro Gln Gln Ala Leu Gly
<210> 1104
<211> 36
<212> PRT
<213> Homo sapiens
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<221> SIGNAL
<222> -19..-1
<400> 1104
Met Asp His Val Val Ile Phe Val Ile Phe Pro Ala Ala Leu Leu Leu
        ··· · -15
                                Cys Trp Gly Gly Leu Ile Pro Leu Cys Ile Ile Tyr Pro Pro Ile Ala
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Asp Thr Val Gly
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524
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 Leu Xaa Phe Asp Ile Tyr Ser Leu Ala Phe Ile His Asp Val
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 Met Leu Phe Gly Leu Arg Gly Met Leu Pro Leu Thr Gln Gln Ala Pro
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· Ile Pro His Leu Arg Cys Lys Leu Ser Val Thr
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 Met Arg Val Cys Met Arg Leu Cys Ala Cys Val Tyr Ala Cys Val Cys
                        -15
                                            -10
 Ala Ser Val Ser Ala Cys Val Tyr Xaa Cys Val Cys Met Xaa Val Arg
 Ala His Leu Cys Val Cys Met Cys Val Cys Met Cys Val His Leu Cys
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 Val Cys Met Cys Val Cys Ala Ser Val Cys Val Cys Met Cys
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 Ala Cys Val Cys Met Cys Val Cys Val Arg Ala Ser Val Cys Val
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                        -15
 Ile Ser Ile Ala Ala Leu Arg
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Met Glu Glu Leu Asp Arg Lys Trp Arg Glu Lys Val Leu Pro Ala Ala
                        -45
Lys Leu Ile Lys Arg Arg Asn Leu Phe Ser Thr Cys Thr Pro Gln Tyr
                    -30
                                        -25
                                                             -20
Gly Thr His Ala Ala Phe Leu Ser Leu His Ala Ser Leu Val Thr Lys
               -15
                                    -10
Ala Phe Ser Ile Asn Ser Trp Glu Trp
                            5
<210> 1110
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Met Val Ser Gly Ala Gln Ala Pro Ser Ser Gln Arg Pro Leu Leu Leu
-25
                    -20
Cys Pro Leu Ser Ser Gly Ser Pro Cys Pro Arg
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Met Ser Cys Leu Leu Arg Ala Tyr Ile Ile Trp Ile Phe Pro Ser Phe
                           -20
                                                -15
Leu Pro Ser Leu Leu Ser Ser Phe Leu Leu Ser Leu Pro Pro Ser Gly
  -10
                        -5
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<211> 67
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Met Phe Gln Leu Leu Ile Leu Cys Gln Met Asn Ser Leu Lys Ile Phe
                       -30
Ser Pro Ile Leu Gly Trp Ser Leu His Phe Val Tyr Cys Phe Leu Cys
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526 -15 -10 Cys Ala Glu Ala Phe Leu Leu Asp Met Ile Pro Phe Met Gln Phe Tyr 1 5 Phe Gly Tyr Leu Cys Leu Trp Gly Ile Thr Leu Lys Ile Phe Ala Gln 20 15 Ser Asn Trp 30 <210> 1113 <211> 54 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -48..-1 <400> 1113 Met Ala Leu Leu Gly Lys Arg Cys Asp Val Pro Thr Asn Gly Cys Gly -40 Pro Asp Arg Xaa Xaa Gly Xaa Asn Pro Gln Xaa Arg Asp His His -30 -25 -20 Gln Xaa Xaa Val Cys Leu Arg Leu His Val Leu Ser Ala Val Gln Thr Glu Arg Arg Gly Asp Gly <210> 1114 <211> 37 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -32..-1 <400> 1114 Met Arg Pro Ala Leu Arg Ser Phe Trp His Ser Ser Gly Gly Pro Pro -25 -20 Pro Ser Ala Thr Leu Ala Leu Leu Ser Ser Asp Ser Val Ala Thr Gly Ser Val Val Ser Arg <210> 1115 <211> 49 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -26..-1 <400> 1115 Met Leu Cys Ala Cys Lys Ala Arg Gly Val Met Leu Leu Leu Phe Ser -20 -15 Gly Trp Leu Val Trp Trp Gly Ser Arg Ser Ser Gln Xaa Leu Arg Met -5 1 Pro Glu Xaa Xaa Val Ser Gly Glu Gly Arg Ser Asp Xaa Xaa Pro His Gly

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527
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Met Ile Ser Ser Ser Leu Ser Gly Arg Val Pro Val Ile Leu Gly Asn
                            -35
                                                -30
Leu Met Cly Val Gly Ala Ala Val Arg Arg Met Gly Phe Ser Leu Ile
                        -20
                                           -15
Leu Pro Thr Ser Pro Ser Pro Ala His Ser Gly Ser Ala Pro Ser Ala
-10
                   -5
                                        1
Gly Pro Arg
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Met Gly Ile Ile Gln Xaa Ile Leu Ala Thr Ser Arg Asp Cys Tyr Ser
                      -40
                                            -35
Phe Lys Lys Pro Ile Pro Lys Lys Pro Thr Met Leu Ala Leu Ala
                   -25
                                    -20
Lys Ile Leu Leu Ile Ser Thr Leu Phe Tyr Ser Leu Leu Ser Gly Ser
                                  -5
              -10
His Gly Lys Xaa Asn Gln Asp Val
       5
                            10
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Met Met Leu Ser Thr Phe Ser Tyr Ala Cys Leu Pro Phe Val Cys Leu
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Leu Leu Arg Asn Val Tyr Ser Asp Leu Leu Pro Asn Arg
<210> 1119
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-15

-10

-20

Val Cys Ile Ser Leu Val Ile Ile Asp Asp Glu His Gly <210> 1120 <211> 18 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -14..-1 <400> 1120 Met Leu Leu Pro Leu Gly Leu Lys Val Leu Gly Leu Gln Ala Arg Gly -5 -10 Thr Thr <210> 1121 <211> 48 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -28..-1 <400> 1121 Met Arg Pro Thr Met Glu Phe His Ser Val Leu Cys Gly Val Thr Pro -25 -20 -15 Thr Leu Leu Val Met Trp Leu Ser Pro Gln Met Ala Ser Ser Pro Ser -10 -5 Gln Ala Pro Gly Met Glu Pro Cys Ala Ser Gly Ile Ser Gln Arg Ala 10 15 20 <210> 1122 <211> 52 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -33..-1 <400> 1122 Met Gly Lys Lys Ile Trp Thr Pro Ser Ser Tyr Pro Met Pro Ser -30 -25 His Lys His Val Ser Leu Cys Leu Leu Thr Val Ala Val Leu Val Leu Thr Phe Lys Ser Leu Ile His Phe Glu Xaa Ile Phe Ala Tyr Glu Ile 1 Gly Val Gln Gly <210> 1123 <211> 31 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -24..-1 <400> 1123 Met Ser Pro Val Leu Cys Phe His Arg Cys Ser Cys Pro Ser Leu Leu

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529
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 Ser Pro Ile Ser Pro Ser Gln Ala Cys Pro Glu Pro Leu Leu Gly
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                                     -15
 Met Cys Val Cys Val Cys Val Cys Val Tyr Arg Leu Phe Ser Ser Ser
             -5
                                 1
 Ser Pro
     10
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 <222> -91..-1
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 Met Lys Ser Thr Val Ser Ser Arg Glu Val Ala Thr Val Asp Lys Met
                         -85
 Lys Arg Arg His Ala Glu Tyr Cys Ala Gln Gly Leu Gln Arg Phe Lys
                                         -65
 Ala Gln Leu Ser Gln Asp Thr Leu Pro Xaa His Pro His Leu Glu Xaa
                                     -50
 Glu Lys Gly Leu Glu Gly Leu Glu Glu Asn Val Pro Leu Lys Gly Glu
                                 -35
                                                     -30
 Lys Pro Gly Glu Gly Pro Glu Ser Pro Lys Lys Arg Arg Arg Val
                             -20
                                                 -15
 Leu Leu Gly Ala Gly Ile Pro Pro Val Ser Ser Ala Pro Arg Arg Gln
 Ser Gln Gln Ala Thr
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 <211> 36
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< <400> 1126
 Met His Asn Ser Cys Arg Pro Val His Leu Phe Phe Phe Phe Xaa
                  -15
                                  -10
 Glu Thr Gly Ser Arg Ser Asn Xaa Trp Leu Glu Xaa Ser Gly Ala Ile
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Ile Ala Asn Ser 15

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<211> 44
<212> PRT
<213> Homo sapiens
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Met Glu Ala Tyr Leu Asn Asp Ser Leu Leu Thr Pro Ser Asp Ser Pro
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Asp Phe Glu Ser Val Gln Ala Gly Pro Xaa Ala Arg Pro Thr Phe Arg
                       -20
Leu Tyr Leu Ser Leu Pro Val Ser Gln Ala Gly Pro
                   -5
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Met Pro Ala Leu Gly Pro Ala Leu Leu Gln Gly Ser Leu Xaa Arg Val
             -10
                                   -5
Gly Pro His Pro Pro Ala Pro Ser Thr Asn Cys Ile His Ser Gln Trp
                           10
                                               .15
His Val Ser Ala Ala Xaa Gly Lys Gly Pro His Leu Arg His Pro Leu
                       25
                                           30
Xaa Gly Xaa Tyr Gln Leu Pro Val Pro Ala Glu Pro Trp Ala Ala Ala
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Gly Gly His Ser Val His
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                                   -10
Cys Cys Ser Ser Tyr
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Ser Phe Leu Gly Asn Trp

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<222> -20..-1

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Met Thr Glu Phe Gly Leu Ser Trp Val Phe Leu Val Ala Ile Phe Lys
-20 -15 -10 -5

Gly Val His Cys Glu Gly Xaa Ile Gly Gly Val Gly Gly Ala

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<211> 47

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<220>

<221> SIGNAL

<222> -24..-1

<400> 1133

Met Trp Ala Ser Ser Pro Trp Pro Ser Ala Trp Ser Cys Cys Leu

-20 -15 -10

Ser Ser Ser Phe Ile Ala Gly Arg Arg Gly Trp Thr Gln Met
-5 1 5

Trp Leu Thr Arg Pro Phe Ser Pro Gln Ala Ser Ser Pro Ser Ala 10 15 20

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<211> 49

<212> PRT

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<220>

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<222> -33..-1

<400> 1134

Met Thr Met Pro Ile Ser Ser Tyr Ser Gln Asn Val Leu Ser Asn Phe
-30 -25 -20

His Asp Gly Tyr Phe Met Leu Ile Ile Leu Ser Ala Ile Leu Leu Asn

~5

-15 -10

Ser Phe Ile Gly Cys Val Ser Phe Tyr His Cys Phe Ser Trp Gly Ser 1 5 10 15

Gly

<210> 1135

<211> 28

.<212> PRT

<213> Homo sapiens

<220>

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<222> -20..-1

<400> 1135

Met Leu Thr His Gly Ala Ser Leu Ser Leu Val Ile Phe Leu Leu Thr
-20 -15 -10 -5

Val Lys His Cys Phe Arg Tyr Arg Val Tyr Lys Thr

<210> 1136

<211> 35

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -22..-1

<400> 1136

Met Ser Ser Val Glu Thr Asp Trp Gly Phe Trp Thr Ser Ile Pro Ile
-20 -15 -10

Leu Pro Leu Ser Ser Gly Arg Gln Leu Pro Leu Pro Thr Arg Glu Trp
-5 1 5 10

Gly Met Trp

<210> 1137

<211> 82

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -33..-1

<400> 1137

Met Phe Ala Ser Pro Arg Arg Trp Ser Ser Xaa Lys Ala Phe Ser Gly
-30 -25 -20

Gln Arg Thr Leu Leu Ser Ala Ile Leu Ser Met Leu Ser Leu Ser Phe
-15 -10 -5

Ser Thr Thr Ser Leu Leu Ser Asn Tyr Trp Phe Val Gly Thr Gln Lys
1 5 10 15

Val Pro Lys Pro Leu Cys Glu Lys Gly Leu Ala Ala Lys Cys Phe Asp 20 25 30

Met Pro Val Ser Leu Asp Gly Asp Thr Asn Thr Ser Thr Gln Glu Val

Val Xaa

<210> 1138

<211> 63

<212> PRT

<213> Homo sapiens

PCT/IB99/00712 533 <220> <221> SIGNAL <222> -16..-1 <400> 1138 Met Pro Ile His Ser Val Phe Leu Cys Ala Pro Ala Leu Val Phe Pro -10 -5 Arg Pro Val Ala Trp Lys Ala Glu Arg Pro Ser Leu Cys Phe Gly Ala 10 Ser Leu Pro Pro Leu Gly Arg Ser Leu Leu Gly Gln Gly Ser Ser Phe 25 Ile Ser Trp Gly Thr Gln Ala Ala Ile Val Glu Leu Xaa Pro His 40 <210> 1139 <211> 80 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -62..-1 <400> 1139 Met Val Tyr Asp Glu Lys Ser Leu Ser Cys Ser His Thr Pro Ala Thr -55 Gln Phe Leu Ser Trp Asp Ala Ser Ser Val Tyr Ser Phe Leu Tyr Ile -35 Leu Ser Ala Arg Val Asn Val Asp Val Xaa Xaa Tyr Ile Arg Val Tyr -25 -20 Ile Leu Ala Cys Val Phe Phe Leu Ser His Pro Leu Phe Xaa Xaa Pro -10 Asn Gly Ser Val Tyr Cys Xaa Arg His Ser Pro Pro Tyr Leu Phe Cys <210> 1140 <211> 38 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -36..-1 <400> 1140 Met Leu Pro Leu Ser Pro Thr Lys Phe Leu Asn Val Phe Leu Gly Leu -30 ~25 Phe Leu Tyr Tyr Leu Gln Leu Val Cys Leu Leu Ile Ile Ser Leu Val -10 Leu Ile Ser Gly Leu Gly <210> 1141 <211> 48 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -29..-1 <400> 1141

Met Asp Lys Val Glu Leu Pro Pro Pro Asp Leu Gly Pro Ser Ser Ala

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534
                -25
                                    -20
Leu Asn Gln Thr Leu Met Leu Leu Arg Glu Val Leu Ala Ser His Asp
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Ser Ser Val Val Pro Leu Asp Ala Arg Gln Ala Asp Phe Val Gln Gly
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Met Gly Gly Thr Ala Gly Trp Ser Ser Gln Asn Thr His Asn Ile Xaa
     -30
                    -25
Val His His Leu Val Trp Leu Trp Phe Val Val Pro Gln Thr Ile Thr
                        -10
Met Ile Thr Pro Lys Ile Thr Glu His Arg Pro Xaa Ile Thr Asp Xaa
Xaa Ile Met Xaa Thr Phe Glu Xaa Leu Gly Glu Leu Pro
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Met Cys Leu Ser Val Ala Leu Tyr Leu Cys Val Cys Val Cys Val Cys
           -15
                               -10
Leu Ile Ala Arg Val Tyr Phe Cys Ile Tyr Val Cys Val Trp
<210> 1144
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<221> SIGNAL
<222> -14..-1
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Met Leu His Leu Leu Phe Gly Leu Phe Pro Val Leu Trp Met Phe Leu
               -10
                                    -5
Val Tyr Phe Phe Leu Ser Ser Phe Phe Phe Phe Phe
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<222> -18..-1
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                                -10
Cys Ala Phe Phe Phe
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<211> 55
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<222> -36..-1
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Met Lys Xaa Asn Asn Leu Arg Arg Gln Ser Pro Ala Leu Arg His Cys
                ~30
                                            -25
Trp Arq Xaa Glu Thr Asp Phe Phe Leu Phe Thr Leu Ile Gly Ala Ser
                                    -10
                   -15
Leu Leu Gln Ser Ala Ser Gly Pro Cys Arg Ile Ser Xaa Xaa Leu Lys
                          · 5
Trp His Ser Lys Gly Thr Leu
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Met Trp Pro Lys Xaa Gly Leu Leu Gly Leu Gly Leu Pro Leu Leu Pro
                   -15
                                       -10
Pro Asn His Pro Ser Val Ala Gln Gly Thr Leu Val Ser Ser His Ser
Gly Ser Gly Ser Glu Gly Arg Val Ala Leu Arg Ser Asp Val His Ser
                            20
Pro Lys Thr Thr Xaa Gln
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Met Tyl Leu Ile Arg Glu Ser His Ala Ser Gly Ser Ser Ser Val Thr
                            -35
                                                -30
        -40
Ser Ser Cys Ser Leu Xaa Ser Xaa Ser Pro Asn Pro Gln Ala Met Ala
    -25
                        -20
                                            -15
Xaa Leu Phe Leu Ser Ala Pro Pro Gln Ala Glu Val Thr Phe Glu Asp
                    -5
                                        1
Val Ala Val Tyr Leu Ser Arg Glu Glu Trp Gly Arg Leu Gly Pro Ala
                                                    20
            10
                                15
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536 Gln Arq Gly Xaa Tyr Arg Asp Val Met Leu Glu Thr Tyr Xaa Asn Xaa 30 35 Val Ser Leu Gly Val Gly Pro Ala Gly Pro Lys Xaa Gly Val Ile Ser 45 Gln Leu Glu Arg Gly Asp Glu Pro Trp Val Leu Asp Val Gln Gly Thr 60 65 Ser Gly Lys Glu His Leu Lys Lys Ser Thr Ala Gln Leu Leu Gly Pro 80 Glu Leu Lys Tyr Lys Glu Leu <210> 1149 <211> 55 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -37..-1 <400> 1149 Met Ile Pro Arg Arg Thr Ser Ala Ser Arg Ala Pro Ser Val Pro Gln -30 -25 Asn Ala Gly Leu Ser Pro Leu Pro Ala Leu Ser Ser Leu Cys Val Ser -15 -10 Trp Gly Thr Ser Ser Thr Val Thr Arg Leu Arg Pro Trp Ile Ser Pro 1 5 Thr Trp Thr Ser Arg Ala Arg 15 <210> 1150 <211> 56 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -14..-1 <400> 1150 Met Val Cys Ile Phe Cys Phe Leu Thr Ser Lys Ala Phe Pro Asn Pro -10 -5 Arg Ser Gln Asp Phe Leu Leu Asp Phe Ser Arg His Xaa Ile Gly Leu 10 Gly Phe Thr Phe Arg Ser Ala Met His Phe Glu Asn Phe Arg Leu Xaa 25 Gly Leu Gly Gln Asp Ser Leu Cys <210> 1151 <211> 25 <212> PRT <213> Homo sapiens <220> <222> -20..-1 Met Xaa Xaa Tyr Xaa Xaa Xaa Gly Phe Cys Ser Val Thr Ser Ser Pro

Met Xaa Xaa Tyr Xaa Xaa Xaa Gly Phe Cys Ser Val Thr Ser Ser Pro
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Leu Ala Ser Ala Gly Arg Thr Thr Arg
1 5

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                               -15
                                                   -10
Leu Ile Ser Val Ala Leu Ser Val Lys Phe His Ile Xaa Gln Gln Val
 -5
Asn Leu Pro Cys Ser Ser
<210> 1153
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            -35
                                  -30
Asn Pro Lys Pro Val Thr Val Pro Ala Phe Leu Xaa Pro Cys Leu Thr
                               -15 .
Ser Phe Ser Cys Xaa Gly Ala Ser Phe Ser Leu Xaa Gly Xaa Arg Arg
Gly Trp Gln His Gly Ser Cys Cys Ser Thr Ile Pro Leu Phe Xaa Thr
                   15
Leu Asn Ser Leu Gly Gln Gly Leu Ile Gly Pro Ala Tyr Ile Gly Ala
               30
                                   35
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  -15
                       -10
Gln Gly Arg
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<211> 31 <212> PRT

<213> Homo sapiens

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<222> -13..-1
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Leu Thr Asp Gln Leu Phe Pro Ala Pro Ala Ser Leu Ile Pro Glu
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Met Asn Gln Asp Phe Asn Pro Glu Ile Glu Ala Ser Pro Gln Val Lys
                -25
                                    -20
Thr Gly Val Phe Leu Phe Ser Ile Ile Gly Ser Phe Gly Phe Pro Gly
            -10
                                -5
Met Cys Asn Cys Lys Asn Pro Ala Arg
   5
                       10
<210> 1160
<211> 24
<212> PRT
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<222> -20..-1
<400> 1160
Met Pro Cys Ser Trp Ser His Ile Val Ser Ser Leu Phe Ser Trp Leu
                    -15
Leu Ser Leu Thr Ser Val Pro Gly
<210> 1161
<211> 31
<212> PRT
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<222> -28..-1
<400> 1161
Met Phe Phe Gly Tyr Ser Glu Asp Ile Tyr Cys Val Ser Gly Pro
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Val Leu Ser Cys Cys Cys Leu Thr Ala Gly Arg Ala Arg Leu Trp
       -10
                            -5
<210> 1162
<211> 58
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<222> -16..-1
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                        -10
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Ser Pro Pro Ala Ser Leu Glu Ala Ser Ser Asn Val Tyr Leu Gln Glu
                5
                                    10
Ser Arg Ala Ala Tyr Ala Ser Val Pro Ala Gly Pro Glu Val Ala Thr
                                25
Gln His Thr Ser Ser Pro Val Thr Pro Met
        35
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Lys Ala Gly Thr
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     -40 --- - -35
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Thr Ser Val Leu Arg Gln Arg Lys Gly Ser Val Arg Lys Gln His Leu
                       -20
                                           -15
Leu Ser Trp Ala Xaa Gln Xaa Gly Arg Xaa Gln Val Val Glu Ile Leu
-10
                   -5
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Gln Ser Glu Lys Gln Thr Xaa Xaa Asp
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                       -30
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Trp Leu Leu Pro Thr Thr Leu Ala Leu His Gly Ser Leu Asp Ala
                           -15
                                               -10
Val Ser Gln Ala Gln Gly Arg Pro Gly His Pro Asp Ala Pro Pro
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Gln Pro Lys Cys Pro
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               -25
                               -20
Arg Leu Cys Phe Leu Ser Val His Phe Arg Leu Arg Trp Gly Asp Ser
               -10
                                  -5
Cys Pro Ser Ser Pro His Arg Glu Thr Phe Pro Ala Gly Pro Val Asn
                          10
Gly Pro Leu Tyr His Pro Arg
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                           -10
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WO 99/53051 542 Ala Gln Asp Val Ser Glu Arg Leu Gly Arg Glu Ala Arg Gly Arg Arg 10 Leu Gly Cys Arg Val Gln Ala Leu Asp Ser Tyr Pro Val Val Asn Leu 20 25 Ile Asn Glu Pro Leu Val Ile Phe Val Cys Ala Thr Xaa Gly Gln Gly 40 Asp Pro Pro Asp Asn Met Lys Asn Phe Trp Arg Phe Ile Phe Arg Lys Asn Leu Pro Ser Thr Ala Arg <210> 1170 <211> 48 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -41..-1 <400> 1170 Met Ser Ser Ile Leu Gly Val Ser Ser Ser Trp Trp Tyr Leu Tyr Tyr -35 -30 Gly Tyr Cys Ile Phe Val Lys Lys Cys Ser Phe Cys Ser Phe Leu Phe -20 -15 Leu Ala Cys Ile Phe Gln Gly Xaa Ser Xaa Xaa Xaa Asn Thr Gln Ser <210> 1171 <211> 51 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -28..-1 <400> 1171 Met Gly Ser Val Leu Gly Leu Cys Ser Met Ala Ser Trp Ile Pro Cys -25 -20 Leu Cys Gly Ser Ala Pro Cys Leu Leu Cys Arg Cys Cys Pro Ser Gly -5 Asn Asn Ser Thr Val Thr Arg Leu Ile Tyr Ala Leu Phe Leu Leu Val Gly Val Trp <210> 1172 <211> 109 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -46..-1 <400> 1172 Met Ser Xaa Xaa Arg Leu Xaa Arg Gln Leu Leu Ser Gln Xaa Arg -40 . -35

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543 10 15 Lys Glu Met Asn Gly Leu Trp Ser Glu Cys Asp Ser Leu Lys Asn Thr 25 Phe Ile Val Trp Xaa Cys Ile Phe Ser Cys Leu Gly Met Gln Leu Xaa 40 45 Ser Ser Xaa Val Ser Asn Val Arg Leu Leu Ser His 55 <210> 1173 <211> 64 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -26..-1 <400> 1173 Met Pro His Pro Leu Ala Thr Ser Ala Phe Leu Arg Ser Ala Phe Pro -20 -15 Phe Val Cys Leu Thr Phe Cys Val Gly Gly Pro Gly Ile Ser Gly -5 Val Tyr Arg Leu Leu Met Ala Asn Ala Thr Arg Arg Glu Ser Glu Val 10 15 Ser Leu Arg Gly Leu Gly Arg Asp Gly Glu Gly Ala Arg Ala Thr Pro 30 <210> 1174 <211> 27 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -23..-1 <400> 1174 Met Thr Val Gly Leu His Ile Leu Arg Asp Ser Leu Met Val Phe Leu -20 -15 Asn Leu Phe Phe Leu Asn Cys Asp Pro His Arg -5 1 <210> 1175 <211> 35 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -21..-1 <400> 1175 Met Val Arg Trp Gly His Pro Pro Met Phe Cys Val Ser Leu Leu Leu -15 -10 His His Ala Tyr Pro Leu Pro Ser Thr Met Ile Val Ser Phe Pro Arg Pro Pro Leu <210> 1176 <211> 93

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  Cys Phe Gly Cys Pro Gly Gly Ala Ser Ser Arg Cys Arg Ser Pro Arg
                     -5
  Gly Arg Gln Ala Ser Arg Val Pro Arg Leu Glu Asn Gly Ala Gln Arg
           10
                                  15
  Val Val Arg Thr Met Val His Leu Val Leu Gln Pro Lys Arg Val Thr
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                              30
  Leu Val His Pro Pro Arg Gly Leu Glu Pro Val Cys Thr Pro Ile Ala
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  Xaa Met Xaa Pro Lys Ser His Gly Leu Arg Ser Ser Leu
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  <211> 47
  <212> PRT
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                                                      - 5
 Tyr Gly Ser Pro Asp Ala Glu Leu Thr Gly Pro Asp Pro Trp Asp
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                                         -5
 Ser
 <210> 1179
 <211> 48
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                                     -30
                                                         -25
 Pro Arg Asp Phe Gly Ile Glu Arg Arg Val Leu Val Cys Phe Phe Asn
                                 -15
                                                      -10
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Ser Val Ser Phe Leu Phe Gly Val Ser Xaa Lys Lys Ser Xaa Gln Trp
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           -10
                                -5
Ala
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<211> 23
<212> PRT
<213> Homo sapiens
<220>
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  -15
                       -10
Ser Thr Phe Xaa Ala Tyr Leu
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<211> 35
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<213> Homo sapiens
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<221> SIGNAL
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Pro Phe Pro Phe Leu Phe Pro Pro Leu Phe Ser Cys Phe Leu Leu
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                                    10
Pro Thr Arg
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<211> 58
<212> PRT
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               -10
                                    -5
Lys Cys Pro Ser Thr Asp Glu Trp Ile Asn Lys Met Trp Tyr Ile Tyr
                           10
Thr Met Glu Tyr Tyr Pro Asp Ile Lys Lys Asn Gly Ile Leu Thr Phe
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Lys Ala Thr Arg Met Asn Arg Lys Thr Leu 40

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<211> 31

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<220>

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<222> -15..-1

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-15 -10

Val Cys Ile Tyr Ile Xaa Val Tyr Val Cys Thr Cys Val Arg Gly 10

<210> 1185

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<220>

<221> SIGNAL

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<400> 1185

Met Gly Val Arg Thr Val Cys His Phe Ile Gln Val Phe Leu Ser Leu -25

-20 -15

Phe Val Phe Phe Trp Leu Val Gly Phe Ser Phe Phe Phe Leu Xaa **--** 5 1

Phe Ser Thr Lys Gln Val Arg Val Glu Gln His Cys Asp Phe Lys Ser 10 15

Thr Pro Xaa Val Glu Ser Ser Ser Thr Val Gly His Ala

30

<210> 1186

<211> 63

<212> PRT

<213> Homo sapiens

25

<220>

<221> SIGNAL

<222> -27..-1

<400> 1186

Met Tyr His Ile Leu Phe Ile His Ser Phe Ile Asp Arg Tyr Leu Ser -20 -15

Cys Phe Tyr Leu Leu Ala Ile Val Ser Asn Ala Val Met Asn Met Gly - 5 1

Val Gln Met Ser Val Leu Ser Pro Cys Phe Ala Phe Val His Ser Ile 10 15

Lys Asn Val Lys Val Leu Cys Phe Leu Leu Phe Phe Leu Phe Gly 30 . .

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<211> 37

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Ser Pro Ser Cys Ala Ala Thr Ile Thr Val Asn Phe Lys Thr Phe Ser
                    1 5
Ser Pro Gln Thr Gly
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<222> -37..-1
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      -35
                                          -25
Ala Glu Arg Ala Asp Pro Ala Pro Val Ser Cys Leu Ser Ala Gly Leu
              -15
                                      -10
Arg Val Cys Cys Ser Gln Arg Ser
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Gln Arg Lys Arg Lys Gln Ser Thr Gln Asp Glu Asp Ala Val Ser Leu
      -45
                        -40
                                         -35
Cys Ser Leu Asp Ile Ser Glu Pro Ser Asn Lys Arg Val Lys Pro Leu
  -30 -25
                          -20
Ser Arg Val Thr Ser Leu Ala Asn Leu Ile Pro Pro Val Lys Ala Xaa
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Pro Leu Lys Arg Phe Ser Gln Thr Leu Gln Arg Ser Ile Ser Phe Arg 10 Ser Glu Ser Arg Pro Asp Ile Leu Ala Pro Arg Pro Trp Ser Arg Asn

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<211> 48

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<220>

<221> SIGNAL

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<400> 1191

Met Val Phe Trp Thr Lys Phe Cys Ile Leu Ile Ser Thr Ala Phe Pro -15 -10

Ser Leu Leu Thr Gln Ile Ile Phe Pro Lys Ser Ile Thr Phe Ala Phe 1 5

Gln Phe Phe Trp Asn Arg Glu Lys Gln Lys Thr Lys Thr Pro Thr Gly 20

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<211> 65

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<220>

<221> SIGNAL

<222> -37..-1

<400> 1192

Met Ala Ser Leu Leu Cys Cys Gly Pro Lys Leu Ala Ala Cys Gly Ile -30 -25

Val Leu Ser Ala Trp Gly Val Ile Met Leu Ile Met Leu Gly Ile Phe -15 -10

Phe Asn Val His Ser Ala Val Leu Ile Glu Asp Val Pro Phe Thr Glu 1

Lys Asp Phe Glu Asn Gly Pro Gln Asn Ile Tyr Asn Leu Tyr Glu His 20

Gly

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<211> 28

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -16..-1

<400> 1193

Met Ser Val Ser Ala Leu Leu Glu Xaa Leu Gln Xaa Ala Ile Pro -10 -5

Arg Xaa Thr Ser Gly Xaa Gln Asp Leu Pro Asn Trp 1 - 5 · . · .__ 10 .___ -....

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Phe Asp Leu Ser Ala Asp Ala Pro Val Phe Gln Gly Leu Ser Leu Val
                   -20
                                        -15
Ser His Ala Pro Gly Glu Ala Leu Ala Arg Ala Pro Arg Thr Ser Cys
Ser Gly Ser Gly Glu Arg Glu Ser Pro Glu Arg Lys Leu Leu Gln Gly
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Pro Met Asp Ile Ser Glu Lys Leu Phe Cys Ser Thr Cys Asp Gln Thr
Phe Gln
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Met Leu Leu His Tyr Leu Lys Leu Lys Gly Asp Gln Trp Lys Leu Ser
                    -30
                                        -25
Ser Val Ser Thr Leu Ile Leu Phe Ile Phe Ile Gly Ser Leu Gln Pro
                -15
                                    -10
Val Pro Thr Arg Phe Lys Arg Phe Ser Cys Leu Xaa His Leu Ser Ser
Arg Asp His Arg Gln Ala Leu Arg
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           -150
                                -145
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Ala Ser Leu Glu Glu Gln Leu Gln Gly Trp Gly Glu Val Met Leu Met
                         -130
                                                -125
Ala Asp Lys Val Leu Arg Trp Glu Arg Ala Trp Phe Pro Pro Ala Ile
    -120
                        -115
                                            -110
Met Gly Val Val Ser Leu Val Phe Leu Ile Ile Tyr Tyr Leu Asp Pro
                    -100
                                        - 95
Ser Val Leu Ser Gly Val Ser Cys Phe Val Met Phe Leu Cys Leu Ala
                -85
                                    -80
Asp Tyr Leu Val Pro Ile Leu Ala Pro Arg Ile Phe Gly Ser Asn Lys
            -70
                                -55
Trp Thr Thr Glu Gln Gln Gln Arg Phe His Glu Ile Cys Ser Asn Leu
        -55
                            -50
Val Lys Thr Arg Arg Arg Ala Val Gly Trp Trp Lys Arg Leu Phe Thr
                        -35
                                            -30
Leu Lys Glu Glu Lys Pro Lys Met Tyr Phe Met Thr Met Ile Val Ser
                    -20
                                        -15
Leu Ala Ala Val Ala Trp Val Gly Gln Gln Val His Asn Leu Leu
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551
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Thr Tyr Leu Ile Val Thr Ser Leu Leu Leu Pro Gly Leu Asn Gln
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His Gly Ile Ile Leu Lys Tyr Ile
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                                            -15
Gly Leu Leu Ala Gly Pro Ala Ala Thr Ser Trp Ser Arg Leu Pro Ala
                    -5
Arg Gly Phe Arg Glu Val Val Glu Thr Gln Glu Gly Lys Thr Thr Ile
                                15
Ile Glu Gly Arg Ile Thr Ala Thr Pro Lys Glu Ser Pro Asn Pro Pro
                            30
Asn Pro Ser Gly Gln Cys Pro Ile Cys Arg Trp Asn Leu Lys His Lys
                        45
Tyr Asn Tyr Asp Asp Val Leu Leu Leu Ser Gln Phe Ile Arg Pro His
Gly Gly Met Leu Pro
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Val Val Ala Trp Thr Phe Ala Ser Asp Ser His Cys Xaa Xaa Val Xaa
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Met Val Xaa Xaa Ser Gln Leu Xaa Asn Pro Pro Leu
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<210> 1202
<211> 48
<212> PRT
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Met Leu Ala Arg Ala Ala Glu Xaa Thr Gly Ala Leu Leu Leu Arg Gly
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                                    -15
                                                        -10
Ser Leu Leu Ala Ser Xaa Arg Ala Xaa Xaa Pro Pro Leu Gly Leu
Xaa Arg Asn Thr Xaa Gly Thr Val Arg Ala Ala Gly Gly Leu Gly
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Ser Ser Ser Glu Val Gly Glu Asn Gly Arg Ser Val Asp Gln Gly Gly
-40 -35 -30

Gly Gly Ser Pro Arg Lys Lys Val Ala Leu Thr Glu Asn Tyr Glu Leu
-25 -20 -15

Val Gly Val Ile Val His Ser Gly Gln Ala His Ala Gly His Tyr Tyr
-10 -5 1

Ser Phe Ile Lys Asp Arg Gly Cys Gly Lys Gly Lys Trp Leu 5 10 15

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<213> Homo sapiens

<220>

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<222> -20..-1

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-20 -15 -10 -5

Pro Ile Leu Ala Ser Pro Val

<210> 1206

<211> 33

<212> PRT

<213> Homo sapiens

<220>

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<222> -17..-1

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                           -10
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Ala Ile Tyr Ile Ser Pro Ser Val Asn Cys Leu Phe Ile Ser Phe Pro
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Ala
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                                   -20
                                                       -15
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His Phe Gly Phe Glu Thr Val Ser Glu Glu Glu Lys Arg Gly Asp Leu
                   10
Thr Ser Val Val Ser Leu Glu Tyr Pro Glu Val Gln Leu Gln Gly Gln
Arg Val Tyr Ala Phe Leu Ser Pro Ile Cys Thr Tyr Gly Ser Glu Gly
Cys Ser Leu Lys
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<222> -35..-1
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Met Glu Asn Leu Pro Phe Pro Leu Lys Leu Leu Ser Ala Ser Ser Leu
                   -30
                                       -25
Asn Thr Pro Ser Ser Thr Pro Trp Val Leu Asp Ile Phe Leu Thr Leu
               -15
                                   -10
Val Phe Ala Leu Gly Phe Phe Leu Leu Leu Pro Tyr Phe Ser Tyr
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Leu Arg Cys Asp Asn Pro Pro
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       -10
Val Arg Cys Ile
   5
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                -40
                                   -35
                                                       -30
Arg Thr Ala Ala Glu Gln Val Gly Cys Lys Gln Arg Ser Phe His Xaa
           -25
                               -20
Pro Cys Pro Leu Leu Phe Pro Gly Ala Cys Phe Pro Cys Pro
                           -5
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                       -10
                                           -5
Pro Val Leu Gly Gly Ser Ser His Ser Ser Ser Xaa Xaa
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      -45
                   -40
                                      -35
Thr Gln Asn Trp Asn Arg Val Glu Ala Gly Asn Ser Tyr Asp Cys Asp
   -30
                       -25
                                          -20
Asp Pro Leu Val Ser Ala Leu Pro Gln Ala Ser Phe Ser Ser Ser Ser
-15
                   -10
                                       -5
Glu Leu Ser Ser Ser His Ser Pro Gly Phe Ala
<210> 1213
<211> 47
<212> PRT
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<222> -31..-1
<400> 1213
Met Met Ser Glu Xaa Ser Gln Asp Leu Val Lys Cys Ala Pro Pro
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-20

555 Xaa Pro Phe Phe Leu Phe Leu Phe Ser Ser Cys Asp Val Pro Val -5 -10 Pro Leu His Leu Leu Gln Trp Leu Gln Ser Phe Leu Arg Pro Arg <210> 1214 <211> 59 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -27..-1 <400> 1214 Met Phe Arg Cys Val Arg Phe Leu Pro Ser Gly Gly Phe Val Val Leu -20 Leu Thr Ser Gly Val Lys Pro Gln Thr Phe Ala Val Ser Val Thr Ala -5 1 Leu Lys Gly Gly Met Pro Gly Val Val His Ser Ser Gly Gly Phe Val 10 Val Leu Leu Thr Ser Gly Ala Xaa Cys Arg Pro 25 <210> 1215 <211> 52 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -30..-1 <400> 1215 Met Arg Val Gly Arg Arg Glu Gly His Pro Leu Phe Pro Asn Val Pro -20 -25 Arg Cys Leu Phe Leu Asn Ala Arg Leu Ala Gly Thr Leu Cys Gln Leu -5 -10 Lys Leu Leu Gln Phe Gly Arg Leu Gly Asn Thr Glu Ser His Leu His 10 Gly Leu Ala Gly 20 <210> 1216 <211> 33 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -31..-1 <400> 1216 Met Tyr Phe Asp Ile Gln Ile Val Ser Asp Val Val Ser Gly Ile Pro -25 -20 Phe Lys Leu Leu Cys Pro Leu Thr Cye Pro His Ris Ser Leu Ser Thr -15 -10 -5 Val <210> 1217 <211> 47 <212> PRT

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Met Leu Phe Ile Phe Ser Asp Ile Asp Trp Lys Met Asp Leu Cys Phe
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Phe Ser Phe Ser Pro Phe Leu Pro Ser Leu Pro Leu Leu Glu Ala Glu
                    -10
                                         -5
Arg Met Arg Val Ser Asp Gln Leu Gln Tyr Thr Thr Gly Xaa Gly
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                                             -25
Val Phe Pro His Leu Thr Val Val Leu Leu Ala Ile Gly Met Phe Phe
                    -15
                                        -10
Thr Ala Trp Phe Phe Val Tyr Glu Val Thr Ser Thr Lys Tyr Thr Arg
               1
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Asp Ile Tyr Lys Glu Leu Leu Ile Ser Leu Val Ala Arq
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Met Lys Gly Ala Leu Lys Leu Ile Ser Thr Asn Phe Ser Leu Cys Gln
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Ser Val Gln Cys Pro Ser Glu Glu Thr Ile Thr Asp Leu Val Ser Val
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Pro Cys Gln Xaa Gly Leu
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Met Thr Ser Gln Pro Val Pro Asn Glu Thr Ile Ile Val Leu Pro Ser
               -65
                                    -60
Asn Val Ile Asn Phe Ser Gln Ala Glu Lys Pro Glu Pro Thr Asn Gln
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557 Gly Gln Asp Ser Leu Lys Lys His Leu His Ala Glu Ile Lys Val Ile -30 Gly Thr Ile Gln Ile Leu Cys Gly Met Met Val Leu Ser Leu Gly Ile -15 -10 -20 Ile Leu Ala Ser Ala Ser Phe Ser Pro Asn Phe Thr Gln Val Thr Ser Thr Leu Leu Asn Ser Ala Tyr Pro Phe Ile Gly Pro Gly 15 <210> 1221 <211> 55 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -40..-1 <400> 1221 Met Val Asp Glu Cys Leu Thr Glu Pro Val Trp Gly Ser Lys Arg Gln -35 -30 Gly Cys Ser Ser Gln Ala Glu Ala Ser Cys Asp Ile Val Ser Ala Ala -15 -20 Cys Lys Cys Gly Ser Ser Gln Ala Ala Ile Asp Cys Glu Thr Ser Ser 5 -5 Cys Ser Glu Asp Phe Pro Val 10 <210> 1222 <211> 31 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -14..-1 <400> 1222 Met Ala Trp Trp Phe Ser Gly Thr Phe Pro Leu Thr His Pro Cys Ser -5 -10 Gly Tyr Gly Ser Leu Met Ala Pro Ser Ser Pro Thr Pro Ser Gly 10 <210> 1223 <211> 78 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -57..-1 <400> 1223 Met Val Ala Lys Asp Tyr Pro Phe 'yyr Leu Thr Val Lys Arg Ala Asn -50 -45 Cys Ser Let Glu Leu Pro Pro Ala Ser Gly Pro Ala Lys Asp Ala Glu -30 -35 Glu Pro Ser Asn Lys Arg Val Lys Pro Leu Ser Arg Val Thr Ser Leu -15 -20 Ala Asn Leu Ile Pro Pro Val Lys Ala Thr Pro Leu Lys Arg Phe Ser 1 ~5 Gln Thr Leu Gln Arg Ser Ile Ser Phe Arg Ser Glu Ser Ala 15

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Met Ser Pro Ala Phe Arg Ala Met Asp Val Glu Pro Arg Ala Lys Gly
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            -25
Val Leu Leu Glu Pro Phe Val His Gln Val Gly Gly His Ser Cys Val
                            -5
Leu Arg Phe Asn Glu Thr Thr Leu Cys Lys Pro Leu Val Pro Arg Glu
                                         15
                   10
His Gln Phe Tyr Glu Thr Leu Pro Ala Glu Met Arg Lys Phe Thr Pro
                                    30
                25
                             .
Gln Tyr Lys Gly Gln Ser Gln Arg Pro Leu Val Ser Trp Pro Ser Leu
                                45
Pro His Phe Pro Trp Ser Phe Pro Leu Trp Pro Gln Gly
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Met Leu Gly Gly Ala Val Ile Ala Gly Arg Pro Leu Gly Arg Trp Glu
                                     -25
Ser Thr Ala Gln Xaa Ile Leu Ala Phe Leu Gln Ser Pro Arg Ala Ile
                                 -10
            -15
 Leu Pro Gly Asn Phe Phe Glu Lys Asn Ala Gln Ile Gln Gly Gly Pro
 Trp Gly Gly Gly Ser Gly Lys Thr Cys Ala Pro Gly Arg Xaa Asp Pro
                                         25
                     20
 Gly Trp Glu Cys Gly Ala Gly Gly Gly Xaa Gly Glu Ala Ala Gly Ser
 Arg Xaa Arg Xaa Ser
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 Met Ser Met Ala Cys Phe Phe His Leu Phe Val Ser Ser Leu Ile Ser
                          -10
 Phe Glu Gln Cys Phe Xaa Met Leu Arg Lys Leu Leu Lys Ile Ile
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                                      -35
Gly Glu Val Trp Phe Pro Gly Trp Gly His Thr Ile Thr His Cys Phe
                                 -20
                                                   -15
               -25
Pro Trp Leu Glu Val Gly Leu Phe Phe Trp Leu His Ala Ala Pro Gly
                              -5
           -10
Arg Ala Ile Ala Leu Pro His Phe Ser Ser Phe Ser Val Gly Gln Xaa
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Val His Leu Val Ser Pro Leu Xaa Xaa Leu Asp Ile Ser Val Glu
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<222> -19..-1
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Met His Leu Leu Gln Glu Glu Leu Leu Leu Leu Pro Arg Gly Leu
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            -15
Cys Gln Val Cys Pro Arg Leu Cys Leu Gln Arg Xaa Val Gly Glu Leu
Gln Xaa Xaa Xaa Pro Asp Val Gly Thr Ala Leu Leu Pro Asp Val Asn
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Arg Thr Ser Cys Thr Thr Trp
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<210> 1229
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<212> PRT
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<222> -28..-1
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Met Cys Leu Ser Cys Ile Gln Gly Ser Phe Phe Val Glu Ile Leu Gln
                            -20
                                            -15
           -25
Leu Val Thr Arg Leu Leu Ser Pro Ser Gln Ser Thr Gln Thr His
                           -5
     -10
 Thr His Thr His Thr His Thr
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 <210> 1230
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 <212> PRT
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<221> SIGNAL

560

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Met Thr Ile Leu Arg Glu Met Xaa Xaa Ser Leu Tyr Val Leu Glu Ala
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    -30
Lys Asp Thr Ala Ile Leu Leu Leu Val Xaa Val Ser Asp Lys Asn Glu
                        -10
Gln Gln Leu Gly Arg Gly Val
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<222> -29..-1
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Met Arg Leu Ser Ser Ser Cys Gly Leu Pro Val Lys Thr Leu Pro Phe
                                    -20
                -25
Ile Cys Cys Asn Leu Tyr Phe Leu Leu Phe Cys Arg Ser Ser Phe Leu
                                -5
            -10
Tyr Phe Gly Tyr Asp Pro Ile Asn Thr Tyr Met Tyr Tyr Asn Val Phe
                                             15
                        10
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 Ser His Ser
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                                                     -55
                                 -60
            -65
 Ser Pro Gly Ser Arg Gly Arg Gly Ser Asp Leu Glu Arg Gly Leu Cys
                                                 -40
                             -45
         -50
 Pro Ala His Pro Gly Ala Pro Pro Leu Pro Arg Pro Pro Asp Arg Leu
                                              -25
                         -30
 Pro His Ser Phe Ser Pro Thr Gly Cys Leu Leu Xaa Pro Leu Leu Val
                                                             -5
                                          -10
                     -15
 Ser Cys Leu Gly Ser Leu Leu Pro Val Thr Gln Thr Leu Gly Ser Phe
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  Ser Ala Gly Pro Cys Phe Arg Thr Leu
  <210> 1233
  خ211> 46
  <212 PRT
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  <220>
  <221> SIGNAL
  <222> -25..-1
  Met His Ser Leu Cys Pro Leu Ser Gln Phe Leu Pro Ile Leu Xaa Ser
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561 -15 -20 -25 Leu Ser Ser Ser Val Pro Ser Arg Ala Gly Ser Ala Phe Pro Ser Ala -5 Leu Gly Pro Leu Tyr Gln Pro Leu Leu Gly Pro Pro Ala Trp <210> 1234 <211> 59 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -44..-1 <400> 1234 Met Arg Thr Gln Val Tyr Glu Gly Leu Cys Lys Asn Tyr Phe Ser Leu -35 Ala Val Leu Gln Arg Asp Arg Ile Lys Leu Leu Phe Phe Asp Ile Leu -20 -15 -25 Val Phe Leu Ser Val Xaa Leu Leu Phe Leu Phe Leu Val Asp Ile -5 -10 Met Ala Asn Xaa Thr Thr Ser Leu Gly Arg Pro 10 <210> 1235 <211> 109 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -45..-1 <400> 1235 Met Gly Gln Phe Thr Ala Ala Met Val Gly Arg Ile Ser Cys Leu Gly -40 -35 Val Trp Lys Leu Pro Arg Val Glu Ser Cys Ser Gln Pro Ala Arg Pro -20 -25 Leu Leu Ser Leu Ala Gln Thr Thr Thr Lys Thr Thr Ala Thr Thr Thr -5 -10 Thr Thr Thr Lys His Ala Thr Cys Ala Leu Ala Tyr Thr Asn Thr Pro 10 Thr Glu Pro Xaa Gln Ala Asp Lys Ala Ser Arg Arg Ala Ser Gly Xaa 30 25 Leu Xaa Xaa Ala Ala Arg His Ile Pro Trp His Gly Ala Thr Ala Ala 45 40 Gln Leu Pro Ala Pro Pro Pro Ser Val Ile Ser Ala Leu <210> 1236 <211> 28 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -18..-1 <400> 1236 Met Leu Ile Phe Ile Ile Ala Ile Leu Phe Pro Asn Ser Gly Ser Cys -15 -10 Phe Ala Phe Ser Cys His Val Ser Phe Phe Phe

10 1 <210> 1237 <211> 58 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -15..-1 <400> 1237 Met Val Arg Cys Ala Cys Phe Pro Phe Pro Phe Ala Phe Cys His -10 Asp Cys Lys Phe Leu Gly Ala Ser Gln Ser Cys Phe Leu Leu Ser Arg 10 Gln Asn Cys Val Ser Thr Gly Xaa Pro Ser Ser Lys Ser Asp Ile Asn 25 Ser Arg Ser Gly Ser Cys Ser Leu Ala Arg 35 40 <210> 1238 <211> 98 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -27..-1 <400> 1238 Met Val Ser Leu Arg Val Gly Ala Ser Pro Phe Arg Phe Pro Leu Ala -20 Pro Leu Xaa Leu Val Phe Ile Ser Leu Leu Pro Ala Pro Phe Phe Pro 1 -5 Thr Leu Ser Phe Pro Cys Cys Cys Val Ser Trp Leu Phe Ser Leu Ser 15 Val Xaa Val Ser Leu Arg Leu Ser Leu Xaa Val Ser Cys Leu Ser Leu 30 Trp Cys Leu Leu Val Leu Phe Leu Ser Pro Thr Leu Tyr Val Ser Asp 50 45 Ser Phe Cys Ser Phe Cys Val Leu Pro Ile Ala Leu Cys Pro Xaa Ala Arg Ser 70 <210> 1239 <211> 72 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -54..-1 <400> 1239 -45 -50

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Gln Phe Thr Pro His Ser Leu Leu
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<212> PRT
<213> Homo sapiens
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<221> SIGNAL
<222> -31..-1
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Met His Phe Pro Ile Gln Ala Thr Phe Xaa Tyr Ser Pro Thr Asp Ser
                 -25
                                    -20
Leu Cys His Leu Tyr Xaa Ser Leu Phe Ser Ser Phe Leu Cys Ser Thr
                   -10
Pro Ala Arg
<210> 1241
<211> 61
<212> PRT
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<222> -36..-1
<400> 1241
Met Ala Leu His Ile Leu Glu Cys Glu Arg Asn Val Cys Phe Val Ala
                       -30
Val Arg Gln Pro Ala His Glu Ser Cys Phe Val Pro Ser Leu Val Thr
                   -15
                                        -10
Gly Ala Leu Gln Gln Ser Gln Thr Gln His Pro Pro Trp Val Cys Pro
Gln Val Gln Gly Ser Tyr Pro Ser Trp Lys Asn Arg Gly
                            20
<210> 1242
<211> 58
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Met Ser Cys Thr His Ser Ser Ser Asn Leu Gly Lys Phe Ser Val His
       -30
                            -25
Arg Glu Tyr Arg Val Leu Xaa Leu Cys Asn Ser Arg Val Ser Phe Thr
                        -10
                                            - 5
Arg Xaa His Val Lys Arg Pro Pro Xaa Arg Leu Cys Val Ser Ser Lys
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Gly bys Leu Phe His Leu Gly Ala Gly Arg
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<213> Homo sapiens

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Leu Phe Gln Lys Gln Xaa Gly Leu Leu Lys Asn Tyr Xaa Ser Pro Gln
                        5
           1
Arg Gln Val Leu Phe Cys Asn Arg
                       20
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Met Ser Tyr Phe Arg Cys Ile Phe Leu Ala Val Leu Ser Lys Ile Ser
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           -15
Trp Ala Val Asn Met Cys Ser Leu Ile Ser Gly Ser Ser
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<211> 39
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Met Leu Cys Ile Met Phe Gly Ile Glu Thr Asn Glu Ile Thr Lys Met
                                    -25
                                                        -20
                -30
Thr Met Ser Phe Leu Leu Phe Leu Ser Ile Ser Leu Ile Thr Leu Tyr
            -15
Tyr Ser Ser Glu Ala Cys Gly
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<210> 1246
<211> 90
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<213> Homo sapiens
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<221> SIGNAL
<222> -39..-1
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Met Cys Gln Ala Arg Ile Ala Leu Asp Arg Cys Asn Leu Arg Thr Ala
                -35
                                    -30
Phe Ile Leu Phe Xaa Leu Ile Leu Ser His Tyr Val Phe Xaa Leu Leu
                                 -15
                                                     -10
            -20
 Ala Pro Phe Leu Thr Arg Ser Ser Pro Ser Trp Asn Ser Tyr Gly Thr
                             1
 Leu Ala Pro Glu Thr Thr Asn Ser Ser Leu Lys Phe Ser Asn Ser Asn
                    15
                                         20
 Asn Gly Ile Ser Asp Leu Ala Xaa Leu Tyr Phe Ser His Val Xaa Lys
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30 Ile Gly Ser Ala Ser Thr Met Gly Tyr Gly 45 50

<210> 1247

<211> 99

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -24..-1

<400> 1247

Met Val Lys Ser Val Ile Phe Leu Ser Phe Trp Gln Gly Met Leu Leu -20 -15

Ala Ile Leu Glu Xaa Cys Gly Ala Ile Pro Lys Ile His Ser Ala Arg

Val Ser Val Gly Glu Gly Thr Val Ala Ala Gly Tyr Gln Asp Phe Ile 15

Ile Cys Val Glu Met Phe Phe Ala Ala Leu Ala Leu Arg His Ala Phe

Thr Tyr Lys Val Tyr Ala Asp Lys Arg Leu Asp Ala Gln Val Pro Thr

50

Tyr Gly Pro Tyr Gly Arg Cys Ala Pro Met Lys Ser Ile Ser Ser Ser 65

Leu Lys Glu

75

<210> 1248

<211> 88

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<221> SIGNAL

<222> -86..-1

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Leu Val Ala Ser Ala Glu Cys Pro Ser Asp Asp Glu Asp Ile Asp Pro -65 -60

Cys Glu Pro Ser Ser Gly Gly Leu Ala Asn Pro Thr Arg Ala Gly Gly -50 -45

Arg Glu Pro Tyr Pro Gly Ser Ala Glu Val Ile Arg Glu Ser Ser Ser -35 -30 -25

Thr Thr Gly Met Val Val Gly Ile Val Ala Ala Ala Leu Cys Ile -20 -15

Leu Ile Leu Leu Xaa Ala Met Tyr <del>-</del> 5

<210> 1249

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Met Ala Trp Thr Pro Leu Trp Pro Thr Leu Leu Thr Leu Cys Ile Gly

-15 -10 Ser Val Val Ser Ser Asp Leu Thr Gln Asp Pro Ala Val Ser Val Ala 1 Leu Gly Gln Arg Val Arg Ile Thr Cys Gln Gly Asp Asn Leu Glu Glu 20 Tyr Phe Ala Ser Trp Tyr Arg Gln Arg Pro Gly Gln Ala Pro Val Leu 35 40 Val Ile Tyr Gly Lys Asn Asn Arg Pro Ser Gly Ile Pro Xaa Arg Xaa 50 55 Ser Gly Ser Lys Ser Gly Asn Thr Ala Leu Leu Thr Ile Xaa Gly Ala 70 65 75 Gln Ala Glu Asp Xaa Ala Asp Tyr Tyr Cys Ser Xaa Arg Asp His Thr 80 85 Asp Asn Arg Trp Val Phe Gly Gly Gly Thr Arg Leu Thr 95 100 <210> 1250 <211> 70 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -20..-1 <400> 1250 Met Glu Ala Glu Phe Tyr Met Xaa Ile Leu Thr Cys Leu Ile Phe Arg -15 -10 -5 Asn Ser Glu Gly Phe Gln Ile Xaa His Val Gln Lys Gln Gln Cys Leu Phe Lys Asn Glu Lys Val Val Gly Ser Cys Asn Arg Thr Ile Gln 15 Asn Gln Gln Trp Met Trp Thr Glu Asp Glu Lys Leu Leu His Val Lys 35 Ser Ala Leu Cys Leu Ala <210> 1251 <211> 19 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -17..-1 <400> 1251 Met Cys Val Cys Ala Cys Ala Leu Cys Val Trp Leu Cys Val Lys Ser ~15 -10 Cys Ser Ile 1 <210> 1252 <211> 34 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -21..-1 <400> 1252 Met Ile Ser Asp Val Gln His Leu Phe Ile Tyr Leu Leu Ala Phe Cys

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Met Pro Ser Leu Glu Lys Cys Leu Tyr Gly Ser Leu Ala His Phe Phe
                                  5
-5
                   1
Phe Phe
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<222> -15..-1
<400> 1253
Met Pro Leu Phe Arg Val Leu Phe Ser Xaa Thr Cys Ala Leu Xaa Gln
        -10
                              -5
Asp Phe Arg Met Gln Pro Cys Pro Pro Thr Pro Lys
           5
                           10
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<211> 30
<212> PRT
<213> Homo sapiens
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Met Trp Tyr Val Glu Met Trp Val Ser Phe Phe Leu Leu Phe Tyr Val
               -20 -15
Leu Leu Phe Arg Asn Leu Tyr Thr His Thr His His Thr Gly
                              1
<210> 1255
<211> 54
<212> PRT
<213> Homo sapiens
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Met Ala Ala Arg Val Gly Ala Phe Leu Lys Asn Ala Trp Asp Lys Glu
                  -25
                                      -20
Pro Val Leu Val Val Ser Phe Val Val Gly Gly Leu Gly Cys Asn Xaa
                                  -5
Ala Pro Ile Glu Pro Leu Leu Gln Val Leu Arg His Asp Gln Gln Gly
                          10
His Ala Leu Gln Leu Xaa
   20
<210> 1256
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<212> PRT
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<222> -23..-1
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<211> 42

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Met Lys Phe Thr His Phe Lys Cys Thr Ile Arg Leu Leu Leu Tyr
          -30
                              -25
Leu Gln Asn Pro Val Thr Ile Thr Ile Leu Phe Leu Ile Val Ser Met
 -15
                          -10
Ala Leu Lys Ile Asn His Ile Pro Lys Gly
             5
<210> 1261
<211> 42
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<222> -14..-1
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Met Ser Cys Met Ser Leu Phe Pro Cys Cys Pro Ala Gln Ser Lys Asn
        -10
                           -5 ·
Tyr Met Leu Leu Phe Ile Ile Leu Leu Pro Thr Gln Phe Leu Tyr
                          10
                                        . 15
Ser Lys Leu Val Thr Ile Cys Cys Cys Phe
                       25
<210> 1262
<211> 26
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -14..-1
<400> 1262
Met Leu Val Cys Cys Thr Ile Asn Ser Ser Phe Ala Leu Gly Ile Ser
              -10
Arg Asn Ala Ile Pro Leu Pro Ala Pro Gly
    5
                          10
<210> 1263
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<212> PRT
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<221> SIGNAL
<2225 -53...-1
<400> 1263
Met Gly Arg Gly Pro Gly Pro Leu Gln Glu Arg Ser Leu Phe Glu Xaa
           -50
                              -45
Lys Arg Gly Ala Pro Pro Ser Ser Asn Ile Glu Asp Phe His Gly Leu
    -35
                          -30
                                              -25
Leu Pro Lys Val Ile Pro Ile Cys Ala Leu Tyr Val Ile Cys Gln Phe
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Ile Leu Ile Arg Ser Gly Val Asn Ile Ser Met Glu Gln Val Thr Val
-5
Val Asp Ala Ser Leu
            15
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<211> 40
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<221> SIGNAL
<222> -13..-1
<400> 1264
Met Leu Tyr Cys Val Val Val His Ser Val Cys Cys Ala Val Tyr
         -10
                                -5
Tyr Phe Val Ile Ile His Thr Ile Glu His Ile Thr Tyr Leu Cys Ile
                       10
His Ser Thr Ile Leu Leu Cys Val
<210> 1265
<211> 37
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<221> SIGNAL
<222> -26..-1
<400> 1265
Met Cys Trp Leu Arg Xaa Trp Gly Gln Ile Leu Leu Pro Val Phe Xaa
                     -20
Ser Leu Phe Leu Ile Gln Leu Leu Ile Ser Phe Ser Glu Asn Gly Phe
                    -5
Ile His Ser Pro Met
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Met Cys Gly Leu Xaa Ile Leu Cys Gly Pro Trp Leu His Ala Ala Pro
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                                    -5
Pro Ser Pro Pro Arg
       5
<210> 1267
<211> 42
<212> PRT
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<222> -33..-1
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<400> 1267 Met Phe His Gly Arg Val Met Ala Met Gly Xaa Leu Thr Lys His Leu -30 -25 -20 Asn Leu Asn Ile Ser Ile Ser Leu Leu Leu Met Leu Xaa Xaa Tyr Trp -10 Ser Cys Trp Ile Lys Ser Pro Pro Xaa Met <210> 1268 <211> 132 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -128..-1 <400> 1268 Met Leu Gly Arg Ser Ser Leu Leu Xaa Trp Lys Xaa Ser Pro Gly Ser -120 Lys Lys Leu Val Val Ala Thr Glu Lys Asn Val Ile Ala Ala Leu Asn -105 -100 Ser Arg Thr Gly Glu Ile Leu Trp Arg His Val Asp Lys Gly Thr Ala -90 -85 Glu Gly Ala Val Asp Ala Met Leu Leu His Gly Gln Asp Val Ile Thr -75 -70 Val Ser Asn Gly Gly Arg Ile Met Arg Ser Trp Glu Thr Asn Ile Gly -60 -55 Gly Leu Asn Trp Glu Ile Thr Leu Asp Ser Gly Ser Phe Gln Ala Leu -40 -35 Gly Leu Val Gly Leu Gln Glu Ser Val Arg Tyr Ile Ala Val Leu Lys -25 -20 Lys Thr Thr Leu Ala Leu His His Leu Ser Ser Gly His Ser Ser Gly -15 -10 Trp Thr Ser Pro <210> 1269 <211> 72 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -57..-1 <400> 1269 Met Ser Thr Thr Tyr Leu Asn Glu Asp Leu Lys Lys Lys Phe Ser Ala -55 -50 Val Ile Glu Gln Val Leu Phe Ala His Leu Ser Pro Leu His Val Trp -35 -30 Leu Gln Leu Arg Ser Leu Cys Glu Xaa Leu Thr Cys Ile Trp Val Arg -20 -15 Phe Asn Phe Leu Ala Ser Ser Gln Ala Cys Ser Lys Cys Asn Ser Ser -5 Phe Leu Ile Met Ser Ser Ser 10 <210> 1270 <211> 80 <212> PRT

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Met Ala Leu Ile Val Leu Gln Leu Thr Phe Gly Ile Gly Tyr Val Thr
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                        -30
Leu Leu Gln Ile His Ser Ile Tyr Ser Gln Leu Ile Ile Leu Asp Leu
Leu Val Pro Val Ile Gly Leu Ile Thr Glu Leu Pro Leu His Ile Arg
Glu Thr Leu Leu Phe Thr Ser Ser Leu Ile Leu Thr Leu Asn Thr Val
                   15
                                        20
Phe Val Leu Ala Val Lys Leu Lys Trp Phe Tyr Tyr Ser Thr Arg Tyr
                                    35
<210> 1271
<211> 54
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<213> Homo sapiens
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<222> -24..-1
<400> 1271
Met Arg Val Ala Gly Ala Ala Lys Leu Val Val Xaa Val Ala Xaa Phe
               -20
                                    -15
Leu Leu Thr Phe Tyr Val Ile Ser Gln Val Phe Glu Ile Lys Met Asp
           - 5
Ala Ser Leu Gly Asn Leu Phe Ala Arg Ser Ala Leu Asp Thr Ala Ala
Arg Ser Thr Lys Pro Pro
<210> 1272
<211> 54
<212> PRT
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<222> -15..-1
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Met His Thr Leu Val Phe Leu Ser Thr Arg Gln Val Leu Gln Cys Gln
                   -10
Pro Ala Ala Cys Gln Ala Leu Pro Leu Leu Pro Arg Glu Leu Phe Pro
                               10
Leu Leu Phe Lys Val Ala Phe Met Xaa Lys Lys Thr Val Val Leu Arg
      20
Xaa Leu Val His Thr Arg
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WO 99/53051 573 <400> 1273 Met Thr Val Val Ile Ser Cys Leu Val Gly Glu Cys Gly Ser Trp Lys <210> 1274 <211> 72 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -47..-1 <400> 1274 Met Cys Thr Leu Thr Asp Thr His Thr His Val Gln Val His Lys Ser -40 Lys Pro Cys Gln Leu Leu Ser Pro Pro Pro Pro Xaa His Gly Pro Leu -25 -20 Leu Leu Pro Ile Phe Gly Leu Leu Val Pro Ser Gln Ile Phe Ser Ser -10 - 5 Leu Leu Asn Ser Leu His Leu Gly Leu Pro Ser Phe Pro Lys Met Pro 5 Leu Met Ile Phe Leu Pro Arg Trp 20 <210> 1275 <211> 78 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -63..-1 <400> 1275 Met Thr Leu Ile Leu Gly Glu Ser Ser Ser Gln Pro Gln Ile Ser Ile -55 -50 Phe Leu Trp Thr Lys Val Lys Asp Leu Phe Ser Leu Met Ile Thr Trp -45 -40 -35 Thr Val Gln Met Lys Leu Thr Ser Met Trp Met Asn Leu Ile Pro Pro -25 -20 Met Lys Gln Ile Leu Xaa Ser Thr Leu Ala Met Lys Ile His Ser Gln -5 -10 Gln Arg Phe Trp Pro Arg Val Arg Val Tyr Ser Arg Ile Tyr <210> 1276 <211> 25 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -19..-1

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Met Asp Ser Val Pro Ala Thr Val Pro Ser Ile Ala Ala Thr Pro Gly
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Asp Pro Glu Leu Val Gly Pro Leu Ser Val Leu Tyr Ala Ala Phe Ile
                           -25
                                               -20
Ala Lys Leu Leu Glu Leu Val Ala Thr Leu Pro Asp Asp Val Gln Pro
                       -10
                                           - 5
Gly Pro Asp Phe Tyr Gly Xaa Xaa Trp Lys Leu Tyr Leu Ser Leu Pro
                                   10
Ser Trp Glu Xaa Phe Val Cys His Phe Leu Met Glu Thr Val Leu Val
           20
                                25
Val Lys Xaa Arg Val Tyr Xaa Val
       35
<210> 1278
<211> 39
<212> PRT
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<222> -18..-1
<400> 1278
Met Ala Ala Tyr Phe Ala Val Trp Ala Ser Val Ala Ser Pro Ala Ser
         -15
                .
                            -10
Ile Cys Cys Gly Xaa Trp Leu Thr Gly Leu Val Arg His Glu Arg Ile
Glu Ala Pro Trp Ala Arg Gly
<210> 1279
<211> 34
<212> PRT
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<222> -29..-1
<400> 1279
Met Lys Thr Gln Phe Leu Ser Trp Gly Lys Phe Ser Phe Cys Phe Gly
               -25
                               -20
Ile Leu Leu Ile Leu Gln Leu Leu Lys Xaa Ser Leu Lys Lys Cys Arg
                               - 5
His Gly
   5
<210> 1280
<2115 40
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<221> SIGNAL
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 Met Leu Pro Ala Val Ala Val Ser Glu Pro Val Val Leu Arg Phe Ile
                     -20
                                         -15
 Leu Pro Ser Ser Trp Asp Cys Arg Cys Ala Pro Pro Leu Leu Thr Gly
                -5
                                     1
 Phe Cys Ile Phe Trp Xaa Glu Thr
 <210> 1281
 <211> 60
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<222> -33..-1
<400> 1281
Met Asp Pro Ala Ala Pro Trp Leu Phe Trp Glu Ala Ala Ala Pro Ala
            -30
                                 -25
Leu Lys Arg Pro Trp Leu Leu Met Val Ala Pro Arg Leu Pro Ala Gly
       -15
                             -10
                                                 -5
Ala Arg Asp Ser Gly Gln Phe Pro Arg Lys Gly Gln Ala Gly Ser Pro
                                        10
Ser Arg Gly Arg Val Arg Lys Leu Gly Gly Ala Val
                20
<210> 1282
<211> 38
<212> PRT
<213> Homo sapiens
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<221> SIGNAL
<222> -31..-1
<400> 1282
Met Lys Met Ser Thr Pro Ser Pro Leu Ser Lys Lys Val Leu Arg Asn
                        -25
                                            -20
Gln Val Ser Arg Leu Xaa Ala Leu Leu Ser Pro Tyr Ala Phe Thr Leu
                    -10
                                        -5
Xaa Arg Leu Ala Ser Gly
<210> 1283
<211> 58
<212> PRT
<213> Homo sapiens
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<221> SIGNAL
<222> -15..-1
<400> 1283
Met Arg Arg Phe Leu Leu Tyr Ala Thr Gln Gln Gly Gln Ala Lys
                   -10
                                        -5
Ala Ile Ala Glu Glu Met Cys Xaa Gln Ala Val Val His Gly Phe Ser
                                10
                                                    15
Ala Asp Leu His Cys Ile Ser Glu Ser Asp Lys Val Ser Val Ile Gln
                            25
Asn Thr Pro Thr Phe Ala Thr Gly Gly Arg
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<400> 1284
Met Leu Ile Asp Ile Trp Ser Met Val Leu Arg Glu Asn Leu Phe Val
    -25
                            -20
Asn Leu Asn Leu Cys Phe Ala Tyr Thr Phe Ala Leu Tyr Ser Cys Pro
                        - 5
Ala Pro Thr Arg Cys Pro Arg Pro Ser
               10
<210> 1285
<211> 73
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<222> -18..-1
<400> 1285
Met Leu Ser Cys Pro Trp Phe Pro Leu Ser Cys Ser Pro Ser Leu Pro
            -15
                               -10
                                                    - 5
Leu Ser Ile Pro Asp Cys Leu Pro Ala Phe Leu Trp Pro Leu Gly Ile
                                           10
Pro Trp Pro Asp Gly Glu Gly Leu Arg Pro Ser Arg Leu Leu Arg Thr
                    20
                                       25
Arg Glu Asn Ile Thr Pro Leu Ser Leu Phe Ala Met Leu Ser Gly Arg
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Glu Gly Ala Pro Leu Leu Val Pro Leu
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<222> -13..-1
<400> 1286
Met Val Val Val Ser Phe Leu Ala Ser Ser Ser Leu Pro Ala Glu Thr
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Pro Lys Gln Gly
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<210> 1.287
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<213> Homo sapiens

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Met Met Gly Ile Phe Leu Val Tyr Val Gly Phe Val Phe Phe Ser Val
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                         -15
Leu Tyr Val Gln Gln Gly Leu Ser Ser Gln Ala
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<400> 1291
Met Ser Leu Gly Leu His Ser Asn Ser Trp Val Leu Asp Pro Ala Leu
                            -15
                                               -10
Leu Leu Thr Cys Leu Thr Phe Pro Ile Tyr Lys Leu Leu Trp Val Arg
                    1
                                     5
Gly Gly Thr Arg Xaa Thr Leu Xaa Ala Leu His Ser Ala Arg Thr
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<210> 1292
<211> 68
<212> PRT
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<400> 1292
Met Ala Ala Asn Ser Ser Gly Gln Gly Phe Gln Asn Lys Asn Arg Val
                   -55
                                       -50
Ala Ile Leu Ala Glu Leu Thr Lys Arg Lys Glu Asn Tyr Leu Cys Arg
               -40
                                   -35
                                                      -30
Thr Ser Leu Gln Gln Ile Ile Leu Glu Leu Gly Ile Asp Thr Ile Met
           -25
                               -20
                                                   -15
Trp Val Xaa Cys Xaa Phe Cys Phe Val Leu Phe Cys Phe Glu Thr Glu
       -10
Ser Arg Pro Val
<210> 1293
<211> 138
<212> PRT
<213> Homo sapiens
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<221> SIGNAL
<222> -35..-1
<400> 1293
Met Ser Ala Gly Ser Ala Thr His Pro Gly Ala Gly Gly Arg Arg Ser
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                -30
                                       -25
Lys Trp Asp Gln Pro Ala Pro Ala Pro Leu Phe Leu Pro Pro Ala
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Ala Pro Gly Glu Val Thr Ser Ser Gly Gly Ser Pro Gly Xaa Thr

-5

WO 99/53051 579 10 Thr Ala Ala Pro Ser Gly Ala Leu Asp Ala Ala Ala Ala Val Ala Ala 20 Lys Ile Asn Ala Met Leu Met Ala Lys Gly Lys Leu Lys Pro Thr Gln 35 Xaa Ala Ser Glu Lys Leu Gln Ala Pro Gly Lys Gly Leu Thr Ser Asn Lys Ser Lys Asp Asp Leu Val Val Ala Glu Val Glu Ile Asn Asp Val Pro Leu Thr Cys Arg Asn Leu Leu Thr Arg Gly Gln Xaa Gln Asp Glu Ile Ser Arg Leu Ser Gly Ala Ala Val Ser 100 <210> 1294 <211> 58 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -21..-1 <400> 1294 Met Ser Pro Leu Asp Gln Ala Val Ile Arg Ala Val Cys Leu Ser Gly -15 Gly Ser Cys Trp Gly Gly Val Arg Cys Leu Val Arg Gly Gly Pro Asn Ile Gly Pro Ala Ala Gln Leu Leu Gly Gly Ile Pro Leu Cys Trp Pro 20 Pro Ala Val Thr Ala Gly Glu Val Lys Leu 30 <210> 1295 <211> 19 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -15..-1 <400> 1295 Met Asn Ser Phe His Phe Ile Xaa Phe Leu Pro Phe Pro Trp Ala Glu -10 Xaa Ala Gln <210> 1296 <211> 35 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -29..-1 <400> 1296 Met Gly Trp His Ser His Ser Ser Gln Gly Val Xaa Ala Met Pro Leu -25 -20 -15 Leu Leu Ser Thr His Thr Trp Thr Asp Thr Ala Leu Ala Phe Ser Thr

-5

His Thr His 5

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 Met Xaa Ala Val Arg Asn Ala Gly Ser Trp Phe Leu Arg Ser Trp Thr
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                             -15
                                                  -10
 Trp Pro Gln Thr Ala Gly Arg Val Val Ala Arg Xaa Pro Ala Gly Thr
    - 5
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                                          5
 Ile Cys Thr
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                    -10
                                         -5
 Phe Ile Ser Pro Ser Ile Gln
 <210> 1299
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Met Tyr Leu Val Cys Thr Thr Cys Thr Trp Cys Val Phe Ser Glu Met
            -50
                                 -45
Phe Val His Gly Leu Asn Ile Thr Gln Leu Val Leu Ser Gln Leu Asp
        -35
                            -30
                                                 -25
Tyr Phe Phe His Ser Asn Leu Thr Asn Leu Val Leu Tyr Phe Leu Val
                        -15
His Leu Leu Phe Ser Leu Ser Leu Phe Met Pro Leu Thr
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<211> 138
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<221> SIGNAL
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<400> 1300
Met Lys Leu Lys Leu Tyr Leu Cys Ile Leu Gly Pro Trp Gly Cys Xaa
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581
 Xaa Lys Val Pro Leu Ile Gly Phe Leu Lys Arg Ile Xaa Xaa Tyr Xaa
                             -55
                                                -50
 Leu Thr Val Leu Lys Pro Xaa Ser Leu Xaa Ser Xaa Ser Ala Gly Leu
                         -40
                                            -35
 Val Pro Ser Glu Asp Ser Lys Lys Glu Ser Val Ser Cys Leu Ser Pro
                     -25
                                        -20
 Arg Phe Trp Trp Leu Gly Ser Leu Xaa Val Thr Trp Leu Ile His
                 -10
                                    -5
 Ala Ser Leu Gln Ser Leu Ser Pro Phe Ser His Ala Ile Phe Ser Cys
                             10
 Val Ser Val Phe Ser Phe Ala Tyr Lys Asp Thr Ser His Ile Glu Leu
                      25
                                            30
 Gly Pro Ala Leu Ile Thr Ser Ser Gln Leu Pro Leu Gln Gly Thr Asn
                    40
 Phe Gln Ile Met Ser His Ser His Val Ala
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 Met Asn Glu Lys Lys Leu Leu Gly Thr Glu Gln Lys Gln Lys Lys
            -30
                        -25
Arg Met Gly Asn Leu Lys Leu Leu Phe Leu Ile Leu Ile Ala
     -15
                           -10
 Gly Tyr Arg
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Met Gly Leu Gln Ser Leu Thr Leu Pro Val Ser Cys Ser Pro Ser Ala
     -25
                           -20
Leu Met Leu Pro Leu Gly Cys Ala Val Arg Thr Arg Met Leu
    -10
                 - 5
<210> 1303
<211> 38
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                       -25
                                          -20
Trp Ile Leu Thr Thr Leu Glu Ser Leu Ala Gly Ser Val Xaa Ser Glu
                   -10
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Gln Asp Leu Ser Ala Tyr
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      -45
                             -40
                                                 -35
 Phe Arg Phe Cys Ser Val Phe Ser Leu Leu Leu Lys Leu Gly Asn Phe
  -30
                         -25
                                             -20
 Tyr Phe Ser Phe Xaa Xaa Cys Leu Phe Leu Xaa Leu Xaa Xaa Ser Glu
 -15
                    -10
                                         -5
 Met Glu Ser His Ser Phe Ser
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Met Glu Asp Val Glu Ala Arg Phe Ala His Leu Leu Gln Pro Ile Arg
                    -60
                                         -55
Asp Leu Thr Lys Asn Trp Glu Val Asp Val Ala Ala Gln Leu Gly Glu
                -45
                                    -40
Tyr Leu Glu Glu Leu Asp Gln Ile Cys Ile Ser Phe Asp Glu Gly Lys
            -30
                                 -25
                                                    -20
Thr Thr Met Asn Phe Ile Glu Ala Ala Leu Leu Ile His Gly Ser Ala
                            -10
                                                -5
Cys Val Tyr Ser Lys Lys Val Glu Tyr Leu Tyr Ser Leu Val Tyr Gln
                    5
                                       10
Ala Leu Asp Phe Ile Ser Gly Lys Arg Arg Ala Lys Gln Leu Ser Ser
                20
                                    25
Val Gln Glu Asp Arg Ala Asn Gly Val Ala Ala Pro Gly Ser Pro Gly
            35
Gly
<210> 1306
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Met Phe Val Ser Tyr Leu Ile Leu Thr Leu Leu His Val Gln Thr Ala
                    -10
Val Leu Ala Arg
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                                        -15
 Tyr Phe Lys Phe Trp Gly Thr Cys Ala Glu Arg Ala Gly Leu Leu His
              -5
                                    1
 Arg Tyr Thr Arg Ala Met Glu Val Cys Cys Thr His Gln Pro Ser Ser
       10
                           15
 Thr Leu Gly Ile Ser Pro Asn Ala Leu Leu Pro Leu
 <210> 1308
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Met Arg Met Gly Thr Arg Ala Ser Pro Pro Leu Cys Met His Leu Ser
           -20
                           -15 .
Ile His Pro Xaa Xaa Cys Ala Cys Ile Cys Pro Ser Ile Gln
                           1
<210> 1309
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<212> PRT
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<400> 1309
Met Tyr Pro Arg Val Trp Gly Cys Phe Gln Leu Leu His Xaa Leu Xaa
                      -30
                                    -25
Xaa Thr Arg Thr Thr Gly Lys Xaa Val Cys Val Cys Val Cys
                   -15
Val Cys Val Cys Val Cys
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Met Ala Ala Val Leu Ala Ala Thr Arg Leu Leu Arg Gly Ser Gly
               -10
                                   -5
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584 Ser Trp Gly Cys Ser Arg Leu Arg Phe Gly Pro Pro Ala Tyr Arg Arg 10 15 Phe Ser Ser Gly Gly Ala Tyr Pro Asn Ile Pro Leu Ser Ser Pro Leu 25 Pro Gly Val Pro Lys Pro Val Phe Ala Thr Val Asp Gly Gln Glu Lys 40 45 Phe Glu Thr Lys Val Thr Thr Leu Asp Asn Gly Leu Arg Val Ala Ser 55 60 Gln Asn Lys Phe Gly Gln Phe Cys Thr Val Gly Ile Leu Ile Asn Ser Gly Ser Arg Tyr 85 <210> 1311 <211> 59 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -25..-1 <400> 1311 Met Tyr Cys Leu Xaa Cys Val Glu Lys Ile Ala Lys Ala Leu Tyr Leu -20 -15 Ser Leu Asn Leu Tyr Phe Ala Asn Ser Leu Tyr Tyr Met Cys Val Cys Ser Tyr Ile Tyr Phe Tyr Leu Xaa Ile Tyr Xaa Tyr Xaa Leu Ile Lys 15 Xaa Xaa Ser Tyr Tyr Val Ala Gln Thr Gly Leu <210> 1312 <211> 36 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -29..-1 <400> 1312 Met Cys Gln Leu Arg Arg Gly Leu Gly Lys Arg Pro Leu Ser Glu Ala -25 -20 Ser Ala Val Phe Leu Thr Ala Val Phe Ser Ser His Ser Trp Leu Val -10 Gly Pro Arg Tyr 5 <210> 1313 <211> 33 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <227> -31..-1 <400> 1313 Met Ser Val Arg Ser Thr Trp Cys Arg Ala Gln Phe Asn Ser Trp Val -25 -20 Ser Leu Leu Thr Phe Cys Leu Ile Asp Leu Ser Asn Val Asp Ser Gly -10 -5

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WO 99/53051
                                       585
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 Gly Arg Arg Ala Arg Lys Leu Leu Pro Ala Pro Arg Ala Ala Pro Arg
       -35
                             -30
                                                 -25
 Thr Ala Pro Asp Tyr Pro Gly Pro Leu Arg Leu Thr Trp Leu Val Ala
                        -15
                                          -10
Ala Gly Leu Glu Gly Arg Val His Leu Ala Asp Thr Ser Ser Gly Arg
                                    5
Lys Thr Trp Pro Gly Cys Gly His Gln Trp Lys Trp Lys Ala Leu Leu
            15
                                 20
Ile Leu Val Arg Ala Phe Pro Ala
       30
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<222> -31..-1
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Met Gly Gly Cys Val Xaa Trp Arg Phe Leu Gly His Ser Ser Ala Leu
                        -25
                                           -20
Arg Thr Val Cys Ser Ser Leu Arg Ser Xaa Arg Pro Cys Trp Cys Asp
                    -10
                                       -5
Gly Leu Arg Leu Arg
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Met Asn Ser Lys Gly Gln Tyr Pro Thr Gln Pro Thr Tyr Pro Val Gln
                       -45
Pro Pro Gly Asn Ser Ser Ile Pro Ser Asp Leu Ala Ser Ser Ser Gly
                   -30
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Ser Thr Leu Tyr Arg Cys Sor Thr Cys Lou Leu Arg Ala Leu Ser Ser -15 Glu Leu Cys Ala Pro Arg Gly Cys His Ser Pro His His Val Ser Arg Ile Ser Trp Thr Leu Ser Val Ser Ser His Gly Pro Val Cys Gly Cys . 20 Trp Ala Phe Arg Phe His Asn Pro His Gly Leu Leu Ser Ser Arg Ser

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30 35 40 45 His Leu Ser Xaa Trp Leu His Ser Ala Gly 50 <210> 1317 <211> 59 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -22..-1 <400> 1317 Met Val Val Ser Ala Phe Ile Tyr Leu Phe Phe Glu Thr Gly Ser -20 -15 -10 Pro Ser Val Ala Gln Ser Gly Val Gln Trp Cys Asp Leu Gly Leu Leu 5 Gln Pro Pro Pro Pro Gly Phe Lys Arg Phe Ser Cys Leu Ser Leu Leu 15 20 Gly Xaa Xaa Asp Cys Arg Arg Ala Pro Pro Gly <210> 1318 <211> 103 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -24..-1 <400> 1318 Met Phe Val Ser Xaa Thr Xaa Phe Phe Phe Xaa Leu Xaa Phe Leu Gly -20 -15 Met Phe Leu Ser Gly Met Val Ala Gln Ile Asp Ala Asn Trp Asn Phe 7 Leu Asp Phe Ala Tyr His Phe Thr Val Phe Val Phe Tyr Phe Gly Ala 15 20 Phe Leu Leu Glu Ala Ala Ala Thr Ser Leu His Asp Leu His Cys Asn 30 35 Thr Thr Ile Thr Xaa Gln Pro Leu Leu Ser Asp Asn Gln Tyr Asn Ile 45 50 Asn Val Ala Ala Ser Ile Phe Ala Phe Met Thr Thr Ala Cys Tyr Gly 65 Cys Ser Leu Gly Leu Ala Leu . 75 <210> 1319 <211> 41 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -26..-1 <400> 1319 Met Ser Ser Glu Ile Phe Xaa Xaa Xaa Ile Ala Tyr Ala Xaa Tyr -20 Leu Leu Val Gly Leu Phe Pro Leu Lys Cys His Xaa Ser Xaa Phe Ser - 5

Lys Xaa Gln Ile Ser Ser Phe Val Glu

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 Cys Ser His Ser Ile Leu Arg Pro Ser Gly Pro Gly Ala Ala Ser Leu
                        5
 Trp Ser Ala Ser Arg Arg Phe Asn Ser Gln Ser Thr Ser Tyr Leu Pro
                    20
                                        25
Gly Tyr Val Xaa Lys Thr Ser Leu Ser Ser Pro Pro Trp Pro Arg
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Met Leu Ile Ala Ala Cys Ile Cys Ser Cys Leu Phe Phe Ser Gln Tyr
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Leu Xaa Xaa Ser Asn Pro Ala Ala
       1
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Met Lys Cys Trp Val Leu Ser Tyr Met Trp Gln Ser Ala Ser Leu Gly
  -15
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Phe Ser Asn Arg Ile Lys Ser Xaa Leu Arg Pro Pro Ala Gly
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Met Ser Val Gly Leu Cys Phe Leu Ile Trp Gln Met Gly Ile Met Leu
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Leu Pro Arg Glu Cys Trp Lys Val Lys Asp Ser Lys Lys Tyr Lys Ser
                                 -45
 Cys Arg Glu Ser Val Leu Pro Ala Gln Ala Cys Thr Gly Glu Ser Pro
                             -30
                                                -25
 Val Leu Ser Gly Val Arg Val Leu Gly Ile Arg Leu Ser Cys Val Leu
                         -15
                                             -10
 Ser His Leu Gln Ala Trp Asp Ser Trp Asp Asn Gln Lys Val Cys Tyr
 Leu Gly Ala Pro Cys Phe Gly Lys Arg Leu Ser Pro Thr Thr Trp Leu
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 Thr Phe Trp Val Gly
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 Met Phe Ala Phe Leu Ala Gly Cys Ser Gly Ser Cys Leu Trp Ser Arg
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                                -5
 His Phe Gly Arg Leu Arg Arg Ala Ala Pro Leu Ser Pro Glu Phe Glu
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 Thr Gly Leu Gly Asn Met Val Glu Pro Gln Trp
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Met Pro Thr Tyr Phe Leu Phe Val Pro His Leu Ile Ser Cys Asn Trp
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Cys Glu Pro Arg Gly Asn Asn Pro Gln Ile Pro Leu Leu Ala Ile His
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Thr Arg Lys Lys Asn Gln His Phe Ile Thr
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                                               -15
Ser Val Ser Leu Xaa Xaa Xaa Xaa Xaa Gly Ser Val Arg Ile Xaa
                -5
                                          1
Leu Ser His Trp Ser Ser Ser Ala Phe Phe Phe Leu Ile Xaa Xaa Xaa
               10
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Xaa Leu Ser His Val Thr Lys Gln Met His Leu <210> 1327 <211> 31 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -14..-1 <400> 1327 Met Leu Thr Cys Leu Cys Gly Cys Phe Ile Val Leu Leu Val Cys Val -10 -5 Leu Lys Cys Val Phe Val Val Ala Ser Asn Gly Leu Phe Pro 10 <210> 1328 <211> 40 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -29..-1 <400> 1328 Met Val Val Ser Phe Ala Val Gln Lys Leu Phe Ser Leu Ile Arg Ser -25 -20 His Leu Ser Ile Leu Ala Phe Val Ala Ile Ala Phe Gly Val Leu Asp -10 Met Lys Ser Leu Pro Thr Pro Gly 5 <210> 1329 <211> 104 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -65..-1 <400> 1329 Met Gly Gly Arg Lys Met Ala Thr Asp Glu Glu Asn Val Tyr Gly Leu -60 -55 Glu Glu Asn Ala Gln Ser Arg Gln Glu Ser Thr Arg Arg Leu Ile Leu -40 Val Gly Arg Thr Gly Ala Gly Lys Ser Ala Thr Gly Asn Ser Ile Leu -30 -25 Gly Gln Arg Arg Phe Phe Ser Arg Leu Gly Ala Thr Ser Val Xaa Arg -15 -10 -5 Ala Cys Thr Thr Xaa Ser Arg Arg Trp Asp Lys Cys His Val Glu Val 5 10 Val Xaa Leu Gly His Xaa Xaa Xaa Gly Lys Cys Pro Arg Gln Ile Leu 20 25 Ala Val Arg Arg Glu Val Thr Ala 35 <210> 1330 <211> 80

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 Pro Gln Ala Leu Ala Leu Thr Xaa Thr Leu Leu Pro Ala Pro Gly Glu
                     -10
                                        -5 .
 His Asp Ser Pro Met Xaa Ile Gly Gln Phe Pro Xaa Asn Pro Pro Ser
                                10
                                                    15
 Glu His Pro Gly Ala Ser Pro Arg Arg Xaa Xaa Thr Gly Trp Xaa Pro
                            25
                                                30
 Gln Ser Trp Asp Arg Arg Val Ser Pro Ala Glu Ala Glu Thr Arg Arg
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                                            45
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Met Gly Val Tyr Thr Cys Pro Ile Phe Val His Tyr Tyr Glu Asn His
                        -35
                                            -30
Gly Pro Thr Pro Ser Phe Xaa Ala Phe Ile Ser Phe His Leu Phe Thr
            -20
Leu Gly Phe Leu Cys Ser Leu Cys Pro His Pro His Gly
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Met Lys Lys Ser Val Ser Cys Cys Ser Ser Leu Trp Val Ser Leu Ser
 -15
                        -10
Lys Asp Glu Asn Ala Glu Met
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                                       -20
Ser Thr Val Leu Leu Ser Gly Ser Pro Arg Ala Val Val Ser Ala Val
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 Cys Trp Cys Leu Ser Phe Pro Thr Ser Ser Phe Thr Glu Ser Val Met
 Arg Ser Leu Gly Glu Cys Pro Arg Lys Arg Trp Gly Gly
    15
                         20
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 Met Xaa Lys Leu Xaa Ser Asn Pro Ser Glu Lys Gly Thr Lys Pro Pro
                -80
                                     -75
 Ser Val Glu Asp Gly Phe Gln Thr Val Pro Leu Ile Thr Pro Leu Glu
            -65
                                 -60
 Val Asn His Leu Gln Leu Pro Ala Pro Glu Lys Val Ile Val Lys Thr
                            -45
 Arg Thr Glu Tyr Gln Pro Glu Gln Lys Asn Lys Gly Lys Phe Arg Val
                        -30
                                             -25
 Pro Lys Ile Ala Glu Phe Thr Val Thr Ile Leu Val Ser Leu Ala Leu
                    -15
                                       -10
Ala Phe Leu Ala Cys Ile Val Phe Leu Val Val Tyr Lys Ala Phe Thr
                1
                                5
Tyr Asp His Ser Cys Pro Glu Asp Ser Ser Xaa Ser Thr Gly
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. <221> SIGNAL
<222> -21..-1
<400> 1339
Met Phe Xaa Ala Ala Gly Val Glu Val Leu Ser Leu Leu Phe Xaa
                        -15
                                            -10
Cys Ile Tyr Trp Gly Gln Tyr Ala Thr Asp Gly Ile Gly Asn Glu Ser
Val Lys Ile Leu Ala Lys Leu Leu Phe Ser Ser Phe Leu Ile Phe
Leu Leu Met
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 Met Leu Thr Gly Arg Phe Leu Gly Gly Ser Gln Gly Phe Phe Leu Ser
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                                             -15
 Phe Leu Ser Phe Phe Phe Phe Phe Phe Leu Raa Phe Phe
 -10
                    -5
 Phe Phe Phe
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Met Phe Ile Xaa Xaa Xaa Met Lys Gln Xaa Phe His Ile Ile Asp Phe
             -25
                                -20
 Val Phe Met Ser Lys Leu Leu Leu Phe Ser Phe Ser Phe Leu Xaa Lys
   -10
                            - 5
Ala Arg Met Xaa Thr Ala Ala Pro Gly
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<211> 37
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<400> 1342
Met Val Thr Pro Val His Ile Leu Thr Ala Val Leu Pro Leu Val Ser
           -15
                               -10
                                                    -5
His Gln Gln Asn His Leu Gly Gly Arg Phe Ala Ser Leu Gly Ser Ser
Gly Ile Arg His Gly
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<211> 19
<212> PRT
<213> Homo sapiens
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Met Leu Ile Leu Mis Leu Ala Thr Leu Leu Asn Leu Phe Ile Ser Ser
-15
                   -10
Asn Ser Phe
<210> 1344
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<212> PRT
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 Met Pro Leu Ala Ser Phe Gly Pro Phe Arg Ser Ser Cys Phe Ala Ala
        -10
                                      - 5
 Arg Ser Ile Ile Trp Lys Ser Gly Arg Gln Gly
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Met Glu Thr Trp Asn Gly Thr Ser Ile Ile Val Ala His Leu Xaa Ser
                      -25
                                    -20
Phe Ser Phe Leu Ser Phe Leu Ser Phe Arg Ser Pro Leu Cys His
                   -10
                                      -5
His Pro Leu Gly
           5
<210> 1346
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Met Gln Phe Leu Ser Leu Ile Phe Ala Ser Cys Ser Ser Thr Thr Pro
         -10
                               -5
Leu Pro Leu Xaa Gln Cys Cys Thr Leu Pro
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<211> 84
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Met Val Thr Ser Lys Ser Arg Gly Pro Xaa Val Gln Thr Leu Gly His
           -50
                              -45
                                   -- 40
Ala why Asn Leu Arg Ser Leu Arg Glu Trp Pro Asp Leu Cys Cys Leu
       -35
                          -30
Arg Leu Phe Val Pro Asp His Thr Val Leu Aia Leu Val Cys His Ser
 -20
                      -15
Ala Ser Ile Ser Val Phe Pro Ser Gln Val Thr Cys Arg Leu Pro Arg
                  1
Thr Gly Ser His Pro Ile Cys Val Ile Ser Gln Gly Ala Phe His Asp
           15
                              20
Pro His Pro Asn
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15

-45

1

-25

-10

WO 99/53051 30 <210> 1348 <211> 53 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -27..-1 <400> 1348 Met Pro Arg Ser Ile Asp Xaa Lys Ala Leu Ile Trp Thr Val Arg Leu -20 Val Val Leu Phe Ala Ser Pro Xaa Val Arg Pro Ala Ser Ser Met Ser -5 Ser Arg Leu Leu Pro Xaa Leu His Tyr Ser Asp Trp Thr Cys Trp 10 Leu Pro Glu Arg Arg 25 <210> 1349 <211> 91 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -54..-1 <400> 1349 Met Thr Ser Leu Leu Thr Thr Pro Ser Pro Arg Glu Glu Leu Met Thr -50 Thr Pro Ile Leu Gln Pro Thr Glu Ala Leu Ser Pro Glu Asp Gly Ala -35 -30 Ser Thr Ala Leu Ile Ala Val Val Ile Thr Val Val Phe Leu Thr Leu -20 -15 Leu Ser Val Val Ile Leu Ile Phe Phe Tyr Leu Tyr Lys Asn Lys Gly -5 Ser Tyr Val Xaa Tyr Glu Pro Thr Glu Gly Glu Pro Ser Ala Ile Val 15 20 Gln Met Glu Xaa Xaa Leu Ala Lys Gly Ser Glu

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<400> 1350

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<222> -23..-1
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Met Ala Gln Leu Ile Met Trp Leu Lys Asn Gln Leu Ile Leu Leu Gly
        -20
                            -15
Ile Phe Arg Gly Ile Arg His Gln Ile Tyr Leu Ile Arg Thr Leu Gln
                            1
Ile Arg Gln Trp
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<211> 91
<212> PRT
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<221> SIGNAL
<222> -30..-1
<400> 1352
Met Gly Pro Val Pro Gly Ala Ala Gly Val Xaa Pro Xaa Xaa Gly
                   -25
                                        -20
Glu Leu Ala Xaa Thr Leu Ser Leu Thr Cys Ser Val Ser Gly Val Ser
                -10
                                    -5
Ile Thr Ser Tyr Tyr Trp Ser Trp Ile Arg Gln Ala Pro Gly Lys Gly
                           10
Pro Glu Trp Ile Gly Xaa Ile Asp His Ser Gly Asp Thr Asp Tyr Asn
                       25
                                            30
Pro Ser Leu Gln Ser Arg Val Thr Leu Ser Val Asp Thr Ser Lys Asn
                  40
Gln Phe Ser Leu Arg Leu Leu Ser Val Ser Ala
               55
<210> 1353
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<222> -36..-1
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Met Trp Phe Gln Thr Arg Ser Cys Gly His His Asp Pro Val Gly Ile
                        -30
                                            -25
Thr Gly Val Thr Lys Val Ile Leu Pro Leu Phe Leu Cys Pro Leu Gly
                                       -10
                   -15
Met Val Glu Thr Ser Phe Gly
               1
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Arg Ile Lys Thr Asn Met Trp Lys His Asn Ile Lys Phe His Gln Leu

Arg Ile Lys Thr Asn Met Trp Lys His Asn Ile Lys Phe His Gln Leu
-60 -55 -50

Pro Tyr Arg Glu Met Glu His Leu Arg Gln Phe Arg Gln Asp Val Thr
-45 -40 -35 -30

Lys Cys Leu Phe Leu Gly Ile Ile Ser Ile Pro Pro Phe Ala Asn Tyr
-25
-20
-15

Leu Val Phe Leu Leu Met Tyr Leu Phe Pro Arg Gln Leu Leu Ile Arg
-10 -5 1

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<222> -19..-1

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Met Tyr Asn Tyr Tyr Phe Leu Ser Leu Pro Ser Phe Leu Cys Thr Cys
-15
-10
-5

Cys Gln Phe Phe Pro His Asp Pro Ile Ser Ser Gln Tyr Ser Ser Pro

1 10

Gln Gly Lys Pro Cys Gln Val Thr Tyr Lys Phe Leu Phe Ile Leu Leu 15 20 25

Gly His Val Tyr Pro Arg Asp Gly Gly 30

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<222> -79..-1

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Met Gln Gly Gly Asn Ser Gly Val Arg Lys Arg Glu Glu Glu Gly Asp
-75 -70 -65

Gly Ala Gly Ala Val Ala Ala Pro Pro Ala Ile Asp Phe Pro Ala Glu
-60 -55 -50

Gly Pro Asp Pro Glu Tyr Asp Glu Ser Asp Val Pro Ala Xaa Ile Gln
-45 -40 -35

Val Leu Lys Glu Pro Leu Gln Gln Pro Thr Phe Pro Phe Ala Val Ala
-30 -25 -20 '

Asn Cln Leu Leu Val Ser Leu Leu Glu His Leu Ser His Val His
*15 -5 1
Glu

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<211> 21

<212> PRT

<213> Homo sapiens

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Met Val Phe Tyr Cys Phe Ala Leu Cys Ile Ile Leu Ile Cys Val Met
     -15
                           -10
Ser Cys Arg His Leu
<210> 1358
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Met Leu Trp Glu Thr Asp Leu Ser Thr Asn Lys Thr Pro Val Ser Cys
          -40
                                -35
Thr Ala Gly Ser Ala Cys Ala Leu Ser Leu Leu Gln Phe Pro Val Leu
      -25
                            -20
                                                -15
Ile Thr Gln Leu Cys Leu Gly Lys Gly Gln Ser Glu Pro Ile Gly Pro
                       -5
Leu Gln Asp Phe Val Ser Leu Glu Ser Thr Ser His Phe Tyr Ser Phe
             10
Phe
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Met Thr Arg Arg Thr Ser Leu Trp Cys Cys Ser Pro Ser Ser Arg
                -15
                                       -10
                                                        -5
Thr Ser Ser Ser Leu Ser Trp Arg Met Gly Ser Gln Ile Arg Pro Ser
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<213> Homo sapiens
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<400> 1360
Met Ala Phe Tyr Leu Trp Cys Phe His Ala Val Phe The Thr Val Cys
           -15
Val Cys Val Arg
       1
<210> 1361
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  <213> Homo sapiens
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  Met Thr Leu Asn Glu His Ala Ala Phe Lys His Leu Phe Asn Lys Ala
             -30
                                 -25
  His Leu Ala Pro Pro Leu Ile His Leu Thr Leu Ser Gly His Ser Thr
                             -10
                                                 - 5
Cys Phe Arg Glu His Arg Val Gly Gly Lys Val Ile Asp Glu Gln His
                                        10
  Pro Lys Ala Glu Glu Ser Phe Leu Val Gln Glu Gly
                 20
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 <222> -26..-1
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 Met Ser Phe Ser Ser Ser Leu Pro Pro Ser Leu Pro Pro Ser Leu Ala
                 -20
                                             -15
 Ser Phe Leu Leu Leu Thr Phe Leu Pro Ser Leu Pro Arg
             -5 ·
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 Met Arg Ala Gln Gly Leu Ser Cys Gly Tyr Pro Ala Arg Pro Leu Gln
                         -40
                                             -35
 Pro Phe Leu Glu His Leu Ala Gly Ser Gly Ile Thr Lys Arg Thr Ala
                    -25
                                         -20
 Pro Gly Cys Ala Pro Leu Arg Trp Val Pro Gln Ile Arg Gly Cys Pro
                 -10
                                    -5
 Leu Thr Arg Leu Ala Gln Arg Gly Ala Asp Thr Arg Thr Arg Glu Asn
                            10
 Leu Phe Tyr Ser Arg Phe Pro Gly Leu Gln Leu Pro Ala Ala Xaa Xaa
                                            30
 Ser Ala Ser Ala Leu Ser Leu Cys Thr Pro Arg Ser Pro Pro Leu Pro
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<213> Homo sapiens

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WO 99/53051
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                            -30
Leu Ala Pro Met Thr Asn Cys His Ser Ile Ser Phe Leu Pro Phe Gln
                                            -10
                        -15
Ala Ser Ile Phe Gly Lys Thr Arg Leu Gln Ser Leu Arg Pro Ser His
Pro Tyr Pro His
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<400> 1365
Met Pro Lys Asp Ala Asp Leu Ala Phe Ser Ala Ser Leu Phe Glu Arg
                -35
                                    -30
Ala Glu Ser Leu Tyr Thr Leu Ile Ser Lys Phe Xaa Ser Cys Xaa Cys
                                -15
Val Ser Thr Leu Ala Tyr Thr Lys Gly Arg Gly
                            1
<210> 1366
<211> 30
<212> PRT
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<222> -28..-1
<400> 1366
Met Phe Val Asn Arg Thr Cys Phe Asn Ser Ser Phe Pro Ile Trp Met
           -25
                          -20
Pro Phe Leu Phe Leu Thr Leu Phe His Cys Leu Gly Arg Arg
                            -5
<210> 1367
<211> 63
<212> PRT
<213> Homo sapiens
<220>
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<222> -37..-1
<400> 1367
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Met Xaa Gly Ser Ser Arg Xaa Xaa Gly Leu Gln Ile Thr Ala Ser Arg -25 -30 Thr Gly Lys Val Tyr Pro Ala Cys His Phe Leu Xaa Ala Val Ser Ala -10 -15 Ser Ser Ser Xaa Ala Cys Leu Trp Tyr Arg Pro Ile Ala Arg Arg Pro 1 Ala Gly Pro Gly Gly Ser Leu Ser Ser Ala Gln Val His Pro Ala 20 15

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<222> -26..-1
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Met Ile Leu Phe Asp His Leu His Cys Ser Ala Ser Gly Val Thr Phe
                        -20
Trp Leu Leu Cys Arg Ile Cys Thr Phe Gly Phe His Gly Phe Ser Lys
Tyr Thr Val Ser Arg Gly Thr Gln Gln Gly Ala Gly Xaa Xaa Xaa Gly
            10
Leu His Gln Asn Trp Glu Gln Trp Arg Gly Leu Val Gly Lys Ser Ser
                            30
Ser Ala Ala Val Val Phe Cys Leu Thr Phe Asp Leu Val Thr Ser Phe
                        45
                                            50
Gln Leu Ala Ser Ala Ile Glu Ser Thr His Phe His Ala Gly Arg Asp
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Gly Ser His Leu
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Met Glu Leu Ser Leu Pro Pro Ser Met Cys Asp Tyr Pro Xaa Phe Cys
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Leu Leu Leu Phe Pro Ala Ser Leu Arg Leu Leu Cys Val His Pro
           -10
                              -5
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Ser Cys Arg Gly Ile Ser Phe Leu Arg Thr Arg
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Met Ser Val Asn Xaa Ile Phe Ile Phe Tyr Phe Ile Leu Leu Leu
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Ile Gln Asp Leu Thr Met Ser Pro Thr Ala Gly Met Gln Trp His Asn
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His Gly Pro Pro Gln Ala Leu Pro Cys Pro Leu Arg Xaa
<210> 1372
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Met Ser Phe Leu Asn Val Asp Ile Thr Asp Cys Leu Tyr Asn Pro Ser
                 -40
                                         -35
Val Cys Pro Val Ala Gln Ser Ser Leu Thr Cys Asp Phe Ile Asp Gly
                -25
                                     -20
Ile Cys Leu Gly Ser Pro Leu Ala Glu Cys Leu Leu Gly Xaa Xaa Xaa
            -10
                                - 5
Xaa Ile Xaa Gly Ile Asn Xaa Xaa Cys Phe Pro Cys Gly Val Lys Cys
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                                            15
Ala Gly Val Val Leu Gly Leu Ser Thr Leu Trp Tyr Val Val
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<211> 49
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Met Lys Val Gly Lys Asp Ser Leu Glu Ser Leu Pro Ser Leu Cys Glu
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        -35
Lys His Ile Gly Pro Ser Gly Leu Phe Thr Phe Leu Ser Pro Ser Phe
                        -15
                                            -10
His Ser Val His Leu Ser Glu Leu Asn Glu Leu Tyr Thr Ile Ala Ala
~5
Gly
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Met Glu Ser Lys Val Leu Ile Ser Ala Ser Leu Leu Arg Ala Ser Gln
      -15
                            -10
Leu Lys Ile Lys Xaa Asn Lys Met Thr Asn Phe Leu Ile Leu
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                -20
Ser Leu Phe Arg Gly Ser His Xaa Xaa Phe Arg Phe Pro Ser Arg Leu
            - 5
Phe Ala Pro Lys Leu Pro Leu Arg Lys Ile Leu Cys Pro Gln Phe Pro
                                            20
Phe Leu Leu Ile Arg Met Ser Pro Gly Asn Ile Trp Asn Gln Lys Asn
                                        35
                    30
Thr Arg Ser Asp Met Val Leu Ala Pro Ser Gly Leu Thr Thr Ala Ala
                                    50
                45
Thr Thr Arg Val Val Tyr Pro His Ser Gly Leu Gly Arg His Val Phe
Val Gly Ile Lys Leu Leu Gly Ile Pro Ala Pro Ser Val Glu Ile Thr
                            80
Ser Cys Met Leu Thr Leu
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Met Lys Ser Asn Leu Thr Leu Leu Thr Cys Leu Xaa Leu Xaa Gly Gly
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                                 -10
            -15
Glu Gly Trp Lys Gly Ala Ala Val Cys Phe Glu Thr Val Glu Gln Phe
                                            . 10
Cys Ser Leu Arg Lys Trp His Val Thr Tyr Leu Xaa Lys Asp Ser Gly
                    20
Leu Cys Gln Gln Gln Glu Lys Leu Tyr Thr Lys Phe Leu Val Cys Ile
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Lys Gly Ala Ser Asn Glu Glu Ile Lys Lys Thr Tyr
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 Met Leu Ala Ser Pro Cys Val Leu Val Gln Gly Ser Gly Xaa Ser Leu
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 Val Arg Thr Pro Trp Cys Pro Glu
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Met Asn Ile Ile Leu Glu Ile Leu Leu Leu Ile Thr Ile Ile Tyr
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                                     -10
Ser Tyr Leu Glu Ser Leu Val Lys Phe Phe Ile Pro Gln Arg Arg Lys
Ser Val Ala Gly Glu Ile Val Leu Ile Thr Gly Ala Gly His
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Met Asp Leu Ile Gly Phe Gly Tyr Ala Ala Leu Val Thr Phe Gly Ser
               -35
                                   -30
Ile Phe Gly Tyr Lys Xaa Arg Gly Gly Val Pro Ser Leu Ile Ala Gly
            -20
                             -15
                                                    -10
Leu Phe Val Gly Cys Leu Ala Gly Tyr Xaa Ala Tyr Arg Val Ser Asn
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Asp Lys Arg Asp Val
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Val Thr Gly Gln Thr Ser Pro Arg Gly Thr Trp Cys Leu Tyr Pro Gly
            1
Phe Cys Arg Ser Val Ala Cys Ala Met Pro Cys Cys Ser His Arg Ser
                        20
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Cys Arg Glu Asp Pro Gly Thr Ser Glu Ser Arg Glu Met Val Arg Val
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Arg Asp His Gly
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<221> SIGNAL

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<210> 1385 <211> 61 <212> PRT <213> Homo sapiens

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           -50
                                    -45
Lys Pro Glu Leu Tyr Glu Val Arg Gln His Gly Arg Ala Val Cys Gly
             -35
                                -30
Gly Glu Asp Asn Ala Ser Pro Gly Glu Gly Leu His Gln Gly Leu Cys
       -20 -15
Leu Pro Gln Arg Val His Cys Ser Leu Leu Pro Ala Pro
                1
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<222> -22..-1
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Met Pro His Ser Phe Val Ser Cys Asn Leu Phe Leu Ser Val Leu Asn
              -15
Phe Leu Phe Leu Leu Ser Phe Ser Thr
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Met Ala Val Phe Leu Gln Lys Arg Lys His Thr Met Arg His His Leu
  -25 -20
Leu Leu Ser Thr Leu Ala Thr Ile Ala Gly Asn Ile Tyr Arg
              · -5
<210> 1388
<211> 47
<212> PRT
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<222> -26..-1
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Met Ala Asp Ser Glu Ala Leu Pro Ser Leu Ala Gly Asp Pro Val Ala
   -25
              ..20
                                       -15
Val Glu Ala Leu Leu Arg Ala Val Phe Gly Val Val Val Asp Glu Ala
-10 -5
                                  1
Ile Gln Lys Gly Thr Ser Val Ser Gln Lys Val Cys Xaa Trp Lys
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<210> 1389

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Met Arg Leu Ala Met Val Gln Leu Val Leu Asn Asn Leu Lys Thr Phe
  -35
                         -30
                                             -25
Tyr Pro Phe Ala Asp His Asp Leu Ala Glu Leu Pro Val Ser Ser Pro
-20
                     -15
                                         -10
Leu Cys His Ala Val Leu Lys Thr Leu Gln Cys Trp Glu Gln Val Leu
Leu Arg Arg Leu Glu Ile His Gly Gly Pro Pro Gln Asn Tyr Ile Ala
        15
                             20
Ser His Thr Ala Xaa Xaa Ser Leu Ser Ala Gly Pro Ala Ile Leu Arg
                         35
His Lys Ala Leu Leu Glu Pro
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<211> 51
<212> PRT
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Met Phe Lys Leu Phe Leu Phe Leu Phe Ile Leu Xaa Tyr Phe Xaa Xaa
                    -15
                                         -10
Tyr Thr Leu Ser Ser Gly Ile Tyr Val Gln Asn Val Gln Val Cys Tyr
                1 .
                                5
Ile Gly Ile His Met Pro Trp Trp Phe Ala Ala Pro Met Asn Leu Ser
        15
                             20
Ser Ala Leu
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<212> PRT
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<222> -21..-1
<400> 1391
Met Ile Tyr Ser Arg Ser Leu Glu Leu Ile Pro Leu Leu Ser Glu Ile
                        -15
Leu Tyr Ala Leu Ala Asn Ile Ser Pro Ile Pro Gln Thr
<210> 1392
<211> 18
<212> PRT
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Met Val His Val Ile Phe Tyr Phe Val Leu Phe Leu Gly Ile Met Thr
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                         -10
Gln Arg
1
<210> 1393
<211> 53
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<221> SIGNAL
<222> -25..-1
<400> 1393
Met His Lys Phe Phe Arg His Phe Tyr Ser Asp Phe Leu Ile Tyr Phe
                    -20
                                         -15
Phe Gln Leu His Ser Cys Cys His Asp Lys Val Thr Ala Xaa Arg Ala
                - 5
Tyr Xaa His Tyr Ser Ser Leu Leu Thr Pro Tyr Leu Ser Gln His Pro
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Cys Pro His Pro Gly
    25
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<212> PRT
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                        -20
                                            -15
Glu Leu Arg Tyr Leu Ser Ala Ala Thr Gly His Pro Ile Ala Thr Pro
Arg Pro Ile Gly Thr Xaa Val Lys Ala Phe Arg Ala His Arg Val Thr
                                15
Ser Glu Lys Leu Cys Arg Ala Gln His Glu Leu His Phe Gln Ala Ala
                            30
Thr Tyr Leu Cys Leu Leu Arg Xaa Ser Gly Asn Met Trp Pro Tyr Ile
                        45
Arg Asn Phe Met Ala Arg Val Ser Ala Arg Trp Arg Ser Leu Leu Ala
                    60
                                         65
Trp Trp Val Ser Ser Cys Pro Ile Ser Leu Glu Gly Arg Ala Gly Ser
                75
His Glu His Gly Glu Tyr Pro Trp Met
            90
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-25 -20 -15 Trp Cys Ser Leu Val Leu Ser Phe Cys Arg Leu His Lys Gln Ser Ser -10 -5 1 Met Thr Val Met Glu Ala Gln Glu Ser Pro Leu Phe Asn Asn Val Lys 10 15 Leu Gln Arg Lys Leu Pro Val 25 <210> 1400 <211> 23 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -14..-1 <400> 1400 Met Arg Leu His Val His Ser Leu Ser Pro Phe Ser Phe Ala Cys Leu -10 -5 Pro Phe Leu Ser Pro Pro Leu 5 <210> 1401 <211> 28 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -26..-1 <400> 1401 Met Leu His Phe Xaa Tyr Met Ile Xaa Val Cys Leu Glu Arg Met Cys -20 -15 Ile Leu Gln Leu Leu Ser Ala Val Leu Tyr Arg Phe -5 <210> 1402 <211> 35 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -30..-1 <400> 1402 Met Ser Ser Glu Pro Pro Pro Pro Pro Pro Pro Pro Thr His Gln Ala -25 -20 Ser Val Gly Leu Leu Asp Thr Pro Leu Gly Ala Val Ser Ala His His -10 Pro Leu Cys 5 <210> 1403 <211> 29 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -20..-1

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 Met Phe Leu Asp His Val Arg Phe Leu Thr Ser Ile Ser Phe Leu Ala
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 Leu Val Leu Trp Asn Val Phe Leu Asn Ser Thr Arg Leu
                 1
 <210> 1404
 <211> 26
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 <222> -19..-1
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Met Arg Glu Lys Pro Gln Pro Ala Leu Leu Thr Ser Ser Glu Leu Pro
            -15
                                    -10
 Ala Leu Ala Ser Gln Ile His Cys Arg Val
                           5
 <210> 1405
 <211> 38
 <212> PRT
<213> Homo sapiens
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<221> SIGNAL
<222> -26..-1
<400> 1405
Met Pro His Asn His Leu Glu Gly Asp Ala Leu Leu Arg Val Pro Val
                        -20
                                           -15
Leu Cys Ile Trp Arg Ala Trp Leu Arg Ala Glu Val Gly Gly Arg Ala
                - -5
                                        1
Pro Leu Pro Gly Arg Met
           10
<210> 1406
<211> 27
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<222> -22..-1
<400> 1406
Met Lys Asn Thr Leu Tyr Tyr Asn Phe Cys Leu Phe Trp Ile Xaa Leu
    -20
                           -15
Pro Pro His Thr Cys Thr His Thr Asp Thr His
   -5
                       1
<210> 1407
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<222> -35..-1
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612 <400> 1407 Met Cys Leu Asn Pro Ala Cys Ser Gly Pro Leu Ser Leu Arg Ser Pro -30 -25 Arg Leu Pro Pro Leu Phe Cys Thr Phe Leu Ser Leu Ser Leu His Pro -15 **-10** . Trp Gly Gly Phe Phe Leu Cys Ala Trp Ile Ser Xaa Phe Leu Pro Trp Val Cys Val Xaa Ala 15 <210> 1408 <211> 101 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -89..-1 <400> 1408 Met Ala His Ser Lys Thr Arg Thr Asn Asp Gly Lys Ile Thr Tyr Pro -85 -80 Pro Gly Val Lys Glu Ile Ser Asp Lys Ile Ser Lys Glu Glu Met Val -70 . -65 Arg Arg Leu Lys Met Val Val Lys Thr Phe Met Asp Met Asp Gln Asp -50 Ser Glu Glu Glu Lys Glu Leu Tyr Leu Asn Leu Ala Leu His Leu Ala -35 -30 Ser Asp Phe Phe Leu Lys His Pro Asp Lys Asp Val Arg Leu Leu Val -20 -15 Ala Cys Cys Leu Ala Asp Ile Phe Arg Ile Tyr Ala Pro Glu Ala Pro -5 Tyr Thr Ser Pro Lys 10 <210> 1409 <211> 26 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -18..-1 <400> 1409 Met Xaa Ser Cys Glu Ile Ala Trp Thr Ala Thr Pro Ser Ser Ala Ala -15 -10 Phe Ala Gln Ala Phe Pro Thr Ala Cys Asn 1 <210> 1410 <211> 46 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -25..-1 <400> 1410 Met Cys His Tyr Leu Trp Lys Lys Leu Tyr Ser Thr Leu Leu Tyr Ile

-20

-15

Leu Ser Arg Ser Ser Gly Arg Gly Lys Asn Leu Ile Thr Ala Val

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613
 Ala Ser Arg Ala Gly Asn Leu Gly Val Trp Thr Glu Lys Gly
                            15
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 <211> 29
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 <213> Homo sapiens -
 <220>
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 <222> -27..-1
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Met Xaa Ser His Arg Leu Phe Gly Cys Phe Pro Ser Asp Leu Ser Arg
                             -20
Met Val Leu Leu Ser Ser Ala Leu Leu Ser Thr Glu Asn
    -10
                         -5
<210> 1412
<211> 47
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<222> -21..-1
<400> 1412
Met Arg Pro Ser His Ser Ser Ala Tyr Leu Cys Leu His Leu Cys Ala
  -20
                        ~15
                                            -10
Phe Ser Thr Glu Gly Trp Met Asn Arg Leu Ser Ser Ser Leu Arg Leu
Ala Pro Leu Pro Leu Tyr Pro Phe Cys Leu Pro Ser Asn Ser Pro
                                 20
<210> 1413
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<400> 1413
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                        -10
Arg Gln Gly Phe Thr Ser Lys Ala Asp Pro Gln Gly Ser Gly Arg Ile
Thr Ala Ala Val Ile Glu His Leu Glu Arg Leu Ala Leu Val Asp Phe
                                25
Gly Ser Arg Glu Ala Val Ala Arg Leu Glu Lys Ala Ile Ala Phe Ala
                            40
Asp Arg Leu Arg Ala Val Asp Thr Asp Gly Val Glu Pro Met Glu Ser
                        55
Val Leu Glu Asp Arg Cys Leu Tyr Leu Arg Ser Asp Asn Val Val Glu
                    70
                                        75.
Gly Asn Cys Ala Asp Glu Leu Leu Gln Asn Ser His Arg Val Val Glu
                85
Glu Tyr Phe Val Ala Pro Pro Gly Asn Ile Ser
            100
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 <222> -81..-1
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                                              -70
 Leu Glu Glu Gly Ser Pro Gly Ser Gly Thr Tyr Thr Arg His Gly Tyr
                     -60
                                          -55
 Ile Phe Ser Ser Leu Xaa Gly Cys Leu Met Lys Ser Ser Glu Asn Gly
                 -45
                                     -40
Ala Leu Pro Val Val Ser Val Val Arg Glu Thr Glu Ser Gln Leu Leu
            -30
                                 -25
Pro Asp Val Gly Ala Ile Val Thr Cys Lys Ser Leu Ala Ser Ile His
        -15
                             -10
Ala Leu Pro
    1
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                                         -50
Thr Met Ala Gln Tyr Gln Ala Ile Ser Lys His Leu Pro Lys Val Cys
                -40
                                     -35
Gln Glu Pro His Leu Pro Arg Gly His Leu Gln Pro Gln Gln His Arg
            -25
                               -20
Leu Leu Val Ala Arg Leu His Met Ala Ser Leu Ala Arg Arg Cys Thr
       -10
                            - 5
Glu Trp Ala Lys Leu His Cys Ser Asp Ala Arg Leu Pro Trp Val Ser
                   10
                                         15
<210> 1416
<211> 35
<212> PRT
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<222> -28..-1
<400> 1416
Met Lys Pro Gln Thr Leu Ala Val Ser Val Thr Val Leu Lys Asp Gly
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                                -20
Val Ala Giy Val Cys Phe Phe Arg Arg Ser Asp Ala Ser Glu Val Ser
       -10
                            -5
Ser Phe Trp
<210> 1417
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WO 99/53051 615 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -43..-1 <400> 1417 Met Val Val Leu Ile Cys Leu Ser Leu Met Ile Ser Asn Thr Glu Leu ~40 -35 Phe Phe Ile Arg Phe Leu Thr Ala Cys Met Pro Ser Phe Glu Lys Cys -20 Leu Phe Leu Ser Phe Ala His Phe Leu Met Gly Arg Thr His Arg . -5 <210> 1418 <211> 36 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -22..-1 <400> 1418 Met Ser Ser Leu Tyr Ile Leu Asp Ile Ser Leu Leu Ser Asp Ile Leu -15 -10 Phe Ala Asn Ile Phe Ser His Ser Trp Asp Val Phe Pro Leu Ser Phe -5 5 Leu Phe Phe Ser <210> 1419 <211> 95 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -84..-1 <400> 1419 Met Gly Gln Gly Ala Arg Gly Trp His Arg Glu Pro Gly Leu Gly Leu -80 -75 Arg His Ser Pro Arg Arg Leu Ser Gly Ala Leu His Leu Glu Ala Gly -65 -60 Cys Asp Arg Asn Ala Thr Thr Val Arg Pro Leu Arg Ala Lys Xaa Gly -45 Asp Ala Leu Pro Glu Glu Ile Arg Glu Pro Ala Leu Arg Asp Ala Gln -30 -25 Trp Val Arg Asp Gln Leu Ala Ser Ser Leu Leu Ile Ile Leu Leu Pro -15 -10 Asn Thr Gln Asp Leu Arg Ile Gln Lys Asp Pro Thr Pro Gly Pro 1 5 <210> 1420

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PCT/IB99/00712 616 <400> 1420 Met Arg Lys Arg Lys Ile Ser Val Cys Gln Gln Thr Trp Ala Leu Leu -45 -40 Cys Lys Asn Phe Leu Lys Lys Trp Arg Met Lys Arg Glu Ser Leu Met -25 Glu Trp Leu Asn Ser Leu Leu Leu Leu Cys Leu Tyr Ile Tyr Pro -20 -10 His Ser His Gln Val Asn Xaa Xaa Ser Ser Leu Leu Thr Met Asp Leu 5 10 Gly Arg Val Asp Xaa Xaa Asn Glu Ser Arg Phe Ser Val Val Tyr Thr 20 25 Pro Val Thr Asn Thr Thr Pro 35 <210> 1421 <211> 33 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -30..-1 <400> 1421 Met Cys Thr Cys Leu Cys Val Cys Leu Tyr Met Tyr Asn Met Gln Phe -25 -20 Leu Xaa Phe Val Phe Val Cys Xaa Leu Leu Lys Cys Met Ser Val Pro -10 Leu <210> 1422 <211> 119 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -31..-1 <400> 1422 Met Ala Ala Ser Ala Ala Ala Glu Leu Gln Ala Ser Gly Gly Pro -25 -20 Arg His Pro Val Cys Leu Leu Val Leu Gly Met Ala Gly Ser Gly Lys -10 -5 Thr Thr Phe Val Gln Arg Leu Thr Gly His Leu His Ala Gln Gly Thr 10 Pro Pro Tyr Val Ile Asn Leu Asp Pro Ala Val His Glu Val Pro Xaa 25 Pro Ala Asn Ile Asp Ile Arg Asp Thr Val Lys Tyr Lys Glu Val Met 45 Lys Gln Tyr Gly Leu Gly Pro Asn Gly Gly Ile Val Thr Ser Leu Asn 60 Leu Phe Xaa Thr Arg Phe Asp Gln Val Met Lys Leu Leu Arg Arg Pro Arg Thr Cys Pro Asn Met Cys 85 <210> 1423 <211> 38 <212> PRT <213> Homo sapiens

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                                        -10
 Tyr Ser Phe Thr His His Leu His Tyr Val Phe Ile Leu Ile Leu Pro
                 1
                                5
 Leu Pro Pro Pro Gln
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<210> 1424
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 <222> -24..-1
 <400> 1424
Met Gly Phe Leu Gly Ser Pro Arg Gln Arg Asn Ser Met Cys Leu Leu
                -20
                        -15
Leu Asp Val Ser Ser Xaa Lys Ser Thr Asp Asn Xaa Xaa Xaa Xaa Xaa
            -5
Leu Ile Ile Tyr Tyr Leu Ile Thr Arg Lys Gly Pro Gly
    10
                       15
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Met Ser Cys Gln Xaa Xaa Leu Ala Xaa Thr Leu Thr Trp Leu Met Ile
                               -35
Arg Gly Arg His Pro Tyr Leu Thr Arg Arg Ser Ala Arg Asn Phe Asn
                       -20
                                               -15
Ile Phe Leu Ala Ala Pro Ser Pro Val Trp Gln Pro Gln Arg Thr Arg
   -10
                      -5
Arg Pro Gln
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<220> <221> SIGNAL

<222> -34..-1

<400> 1426

Met Cys Pro Wie Trp Lou Pro Cys Trp Thr Ala Gln Thr Glu His Leu -30 -25 -20 Asp Arg Tyr Arg Lys Phe His Gln Met Ala Leu Xaa Pro Gly Thr Ser -15 -10 - 5 Arg Ala Gln Ala Leu Leu Tyr Asn Glu Val Leu Glu Arg Phe Met Phe Thr Arg Leu

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618
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 <400> 1427
 Met Asn Val Met Lys Arg Ile Cys Thr Phe Leu Leu Pro Ser His Ser
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 Thr Ser Gly Pro Leu Cys Cys Ser Asn Ala His Leu Pro Ala Thr Ser
 Ser Thr Leu Lys His Cys Arg Ala Trp Arg Glu Ala
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Met Val Val Phe Gly Tyr Glu Ala Gly Thr Lys Pro Arg Asp Ser Gly
                        -115
                                             -110
Val Val Pro Val Gly Thr Glu Glu Ala Pro Lys Val Phe Lys Met Ala
                    -100
                                         -95
Ala Ser Met His Gly Gln Pro Ser Pro Ser Leu Glu Asp Ala Lys Leu
                -85
                                     -80
Arg Arg Pro Met Val Ile Glu Ile Glu Lys Asn Phe Asp Tyr Leu
            -70
                                 -65
Arg Lys Glu Met Thr Gln Asn Ile Tyr Gln Met Ala Thr Phe Gly Thr
                            -50
                                                 -45
Thr Ala Gly Phe Ser Gly Ile Phe Ser Asn Phe Leu Phe Arg Arg Cys
                        -35
                                            -30
Phe Lys Val Lys His Asp Ala Leu Lys Thr Tyr Ala Ser Leu Ala Thr
                    -20
                                        -15
Leu Pro Phe Leu Ser Thr Val Val Thr Asp Lys Leu Phe Val Ile Asp
                -5
                                    7
Ala Leu Tyr Ser Asp Asn Ile Ser Lys Glu Asn Cys Val Phe Arg Ser
                            15
Ser Leu Ile Gly Ile Val Cys Gly Val Phe Tyr Pro Ser Ser Xaa Ala
Phe Thr
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<210> 1429
<211> 63
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<222> -38..-1 <400> 1429

Met Ala Glu Ile Thr Asn Ile Arg Pro Ser Phe Asp Val Ser Pro Val

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-30
  Val Ala Gly Leu Ile Gly Ala Ser Val Leu Val Val Cys Val Ser Val
                                                     -25
                             -15
                                                 -10
  Thr Val Phe Val Trp Ser Cys Cys Xaa Gln Gln Ala Glu Lys Lys His
                         1
                                       5
 Lys Asn Pro Pro Tyr Lys Phe Ile His Met Leu Lys Gly Xaa Ser
                 15
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 Met Val Ile Leu Thr Met Leu Ile Leu Leu Ile His Glu His Gly Ile
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                                         -5
 Phe Phe Ser Leu Val Cys Val Leu Phe
             5
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Met Phe Ser His Asn His Ser Tyr Thr Tyr Thr Pro Gln His Ser Pro
                -25 -20
Leu Thr His Thr His Thr Cys Thr Pro Pro Ser Thr Ala His Pro Arg
Gly
<210> 1432
<211> 22
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Met Phe Xaa Met Ile Leu Leu Cys Phe Leu Ala Val Ser Asn Phe Asn
                   -10
                                       -5
Lys Leu Leu Trp Gly Xaa
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620 <400> 1433 Met Phe Leu Ile Leu Gly Lys Phe Ser Arg Val Met Gly Leu Pro Leu -20 -15 Ala Cys Phe Ser Leu Phe Gly Xaa Leu Pro Gln Gly Leu Leu Ile <210> 1434 <211> 30 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -19..-1 <400> 1434 Met Val Ala Leu Gly Gln Leu Ala Xaa Leu Pro Gly Xaa Xaa His Gly -15 -10 Gly Leu Ser Ala Val Thr Val Val Leu Pro Ile Leu Leu Cys 5 <210> 1435 <211> 22 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -15..-1 <400> 1435 Met Pro Val Ser Phe Val Cys Leu Leu Phe Arg Asn Val Tyr Ser Asn -10 Leu Leu Pro Ser Phe Phe <210> 1436 <211> 64 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -27..-1 Met Gly Ser Gly Gly Asp Ser Leu Leu Gly Gly Arg Gly Ser Leu Pro -25 -20 -15 Leu Leu Leu Pro Ala His His Gly Arg His Gly Ser Gly Leu Pro Ala -5 Pro Asp Pro Ser Pro Pro Pro Gly Pro Ala Val Pro Gly Pro Trp Pro 15 Cys Gln Asp Glu Leu Pro Ser Leu Arg Pro Ala Thr Ser His His Phe 30 <210> 1437 <211> 43 <212> PRT <213> Homo sapiens <220> <221> SIGNAL

<222> -25..-1

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                                         -15
Ser Gly Phe Ser Phe Gln Val Ser Gly Trp Gly Trp Gly Glu Arg Val
                -5
 Asp Asp Phe Leu Phe Ser Ser Gly Ile Asp Gly
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<211> 34
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Met Arg His His Val Arg Xaa Pro Ala Leu Ser Ser Leu Ala His His
                        -15
Pro Arg Thr Ser Gly Gln Lys Arg Glu Pro Ile Ala Pro Ala Gln Leu
                                    5
Ser Pro
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Met Leu Ile Leu Asn Gly Phe Arg Gly His Ala Thr Asp Ser Val Lys
            -70
                                -65
Asn Ser Met Glu Ser Met Asn Thr Asp Met Val Ile Ile Pro Gly Gly
                            -50
Leu Thr Ser Gln Leu Gln Val Leu Asp Val Val Val Tyr Lys Pro Leu
                        -35
                                             -30
Asn Asp Ser Val Arg Ala Gln Tyr Ser Asn Trp Leu Leu Ala Gly Asn
                    -20
                                        -15
Leu Ala Leu Ser Pro Thr Gly Asn Ala Lys Lys Pro Pro Leu Gly Leu
                -5
Phe Leu Glu Trp Val Met Val Ala Trp Asn Ser Ile Ser Ser Glu Ser
                            15
Ile Val Gln Gly Xaa Lys Glu Val Pro Tyr Leu Xaa Gln Leu Gly Gly
                        30
Gly Arg Arg
<210> 1440
<211> 34
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<222> -25..-1
<400> 1440
Met Ile Cys Thr Thr Val Tyr Ile Thr Met Ala Pro Tyr Cys Leu Ser
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WO 99/53051 Asn Cys Leu Leu Xaa Xaa Ser Trp Gly Leu His Leu Tyr Arg Phe Leu -5 1 Ala Pro <210> 1441 <211> 16 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -14..-1 <400> 1441 Met Val Ser Leu Cys Val Ala Ala Leu Phe Pro Leu Gln Ala Tyr Gly -10 <210> 1442 <211> 28 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -24..-1 <400> 1442 Met Leu Ser Ile Phe Ser Phe Phe Cys Arg Pro Phe Val Tyr Leu Leu -20 -15 Leu Arg Asn Leu Xaa Ser Tyr Ser Leu Pro Thr Thr <210> 1443 <211> 94 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -77..-1 <400> 1443 Met Phe Pro Val Ser Ser Gly Cys Phe Gln Glu Gln Glu Thr Asn -70 Lys Ser Leu Pro Arg Ser Ala Ser Thr Pro Glu Thr Arg Thr Lys Phe -55 -50 Thr Gln Asp Asn Leu Cys Xaa Ala Gln Arg Glu Arg Leu Asp Ser Ala -40 -35 Asn Leu Trp Val Leu Val Asp Cys Ile Leu Arg Asp Thr Ser Glu Asp -25 -20 Leu Gly Leu Gln Cys Asp Ala Val Asn Leu Ala Phe Gly Arg Arg Cys -5 Glu Glu Leu Glu Asp Ala Arg His Lys Leu Gln Xaa His Leu

<210> 1444 <211> 20 <212> PRT <213> Homo sapiens <220> <221> SIGNAL

WO 99/53051 623 <222> -15..-1 <400> 1444 Met Pro Leu Val His Ser Phe Leu Trp Leu Ser Ser Ile Leu Tyr Ile -15 -10 Tyr His Leu Arq <210> 1445 <211> 56 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -24..-1 <400> 1445 Met Ile Ser Asn Gly Lys Phe Phe Cys Phe Phe Xaa Val Phe Xaa Phe -20 -15 Xaa Phe Leu Xaa Arg Xaa Leu Xaa Xaa Yaa Pro Arg Leu Glu Cys Asn 1 Gly Lys Xaa Ser Ala His Xaa Asn Leu Arg Leu Leu Ser Xaa Ser Asn 15 Ser Leu Ala Ser Ala Pro Arg Gly <210> 1446 <211> 101 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -90..-1 <400> 1446 Met Glu Asp Ser Ala Ser Ala Ser Leu Ser Ser Ala Ala Ala Thr Gly -85 -80 Thr Ser Thr Ser Thr Pro Ala Ala Pro Thr Ala Arg Lys Gln Leu Asp -70 -65 Lys Glu Gln Val Arg Lys Ala Val Asp Ala Leu Leu Thr His Cys Lys -55 -50 -45 Ser Arg Lys Asn Asn Tyr Gly Leu Leu Leu Asn Glu Asn Glu Ser Leu -40 -35 -30 Phe Leu Met Val Val Leu Trp Lys Ile Pro Ser Lys Glu Leu Arg Val -20 -15 Arg Leu Thr Leu Pro His Ser Ile Arg Ser Asp Ser Glu Asp Ile Cys -10 -5 Xaa Phe Thr Lys Asp <210> 1447 <211> 59 <212> PRT <213> Homo sapiens

<220>

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<222> -29..-1

<400> 1447

Met Asn Ala Glu Gly Ala Ser Pro Gly Lys Glu Thr Asn Thr Gly Thr

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-25
                                      -,20
  Leu Ile Glu Leu Asn Leu Xaa Ser Pro Val Ala Leu Gln Trp Pro Leu
                                                         -15
             -10
                         -5
  Ser Ser Pro Ser Cys Leu Arg Ile Leu Ser Asn Lys Val Pro Arg Asn
                         10
  Leu Arg Trp Gln Lys His Tyr Ser Thr His Gln
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  <211> 81
  <212> PRT
  <213> Homo sapiens
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 <222> -63..-1
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 Met Leu Gly Leu Asp Glu Leu Gly Arg Ser Gly Cys Gly His Cys Thr
                                 -55
 Gln Ala Asp Leu Arg Phe Gly Asp Ala Ala Gly Xaa Glu Pro Arg Xaa
        -45
                             -40
                                                 -35
 Arg Xaa Thr His Arg Asn Thr Ala Ala Ala Arg Val Pro Pro Pro
                        -25
 Arg Val Met Ala Ala Ala Ala Leu Arg Ala Pro Ala Gln Ser Ser
                                             -20
                    -10
                                         -5
 Val Thr Phe Glu Asp Val Ala Val Asn Phe Ser Leu Glu Glu Trp Ser
             5
                                10
 Leu
 <210> 1449
 <211> 49
 <212> PRT
 <213> Homo sapiens
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<222> -26..-1
<400> 1449
Met Ser Ala Leu Lys Asp Phe Arg Glu Phe Leu Asn Trp Trp Gly Asn
                        -20
Leu Ser Phe His Leu Gln Glu Ala His Gly Ser Glu Ile Ala Glu Met
                  -5
Gly Ala Gly Ile Leu Glu Glu Lys Asn Tyr Gly Gln Gln Xaa His Cys
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Asn
<210> 1450
<211> 36
<212> PRT
<213> Homo sapiens
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<221> SIGNAL
<222> -30..-1
<400> 1450
Met Ser Leu Pro Pro Phe Phe His Pro Ser Pro Ala Pro Ser Leu Ala
                   -25
                               -20
Pro Pro Pro Ser Leu Phe Leu Ser Leu Pro Pro Ser Leu Ser Pro Pro
                                   -5
Leu Pro Ala Arg
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 <212> PRT
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 <221> SIGNAL
 <222> -13..-1
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 Met Phe Phe Leu Cys Gly Phe Leu Tyr Leu Cys Phe Ile Ser Phe Phe
            -10
 Phe Phe
   5
 <210> 1452
 <211> 51
 <212> PRT
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<220>
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<222> -42..-1
<400> 1452
Met Lys Ala Gly Pro Cys Ser Cys Gln Glu Gly Gly Arg Gln Trp Ala
        -40
                             -35
His Gly Ser Val Pro Leu Gln Pro Thr Ala Arg Leu Ala Ala Leu Gly
   -25
                        -20
                                           -15
Ile Phe Leu Cys Pro Gly Glu Thr Leu Ser Ala Ser Leu His Trp Asn
Pro Ile Gly
<210> 1453
<211> 53
<212> PRT
<213> Homo sapiens
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<221> SIGNAL
<222> -23..-1
<400> 1453
Met Leu Ser Gln Ser Phe Gln Lys Asn Lys Thr Asn Leu Leu Cys Leu
                                -15
Thr Phe Gln Arg Cys Gln Ser Tyr Asn Trp Leu Asn Ile Phe Glu Ala
                            1
Thr Tyr Met Thr Thr Leu Phe Ile Ser Val Ile Xaa Thr Asn Phe Leu
Lys Arg Tyr Leu Leu
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<222> -30..-1

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<222> SIGNAL <222> -31..-1

<400> 1456

Met His Glu Tyr Leu Pro Arg Asn Phe His Asp Phe Asn Ser Pro Asn
-30 -25 -20

Ser Lys Leu Gly Met Gly Met Gly Phe Phe Ser Gly Val Lys Ser Trp
-15 -10 -5 1

<210> 1457
<211> 83
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<222> -36..-1

<400> 1457

<210> 1458

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627
<211> 24
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Met Val Ile Ser Ala Gly Ala Leu Leu Trp Met Ala Trp Asp Gly Gln
     -15
Leu Ser Arg Pro Glu Gly Ala Arg
<210> 1459
<211> 31
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<222> -18..-1
<400> 1459
Met Val His Cys Asn Leu Glu Leu Leu Gly Ser Ser Tyr Asn Pro Ile
     -15
                               -10
Ser Ala Ser Pro Val Ala Arg Thr Ile Ser Cys Pro Ala Ile Val
                       5
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<211> 127
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Met Leu Gly Ser Gly Phe Lys Ala Glu Arg Leu Arg Val Asn Leu Arg
                    -80
Leu Val Ile Asn Arg Leu Lys Leu Leu Glu Lys Lys Lys Thr Glu Leu
                           -65
Ala Gln Lys Ala Arg Lys Glu Ile Ala Asp Tyr Leu Ala Ala Gly Lys
                       -50
                                           -45
Asp Glu Arg Ala Arg Ile Arg Val Glu His Ile Ile Arg Glu Asp Tyr
                   -35
                                       -30
Leu Val Glu Ala Met Glu Ile Leu Glu Leu Tyr Cys Asp Leu Leu
                                   -15
Ala Arg Phe Gly Leu Ile Gln Ser Met Lys Glu Leu Asp Ser Gly Leu
Ala Glu Ser Val Ser Thr Leu Ile Trp Ala Ala Pro Arg Leu Gln Ser
                      15
Glu Val Ala Glu Leu Lys Ile Val Ala Asp Gln Leu Cys Pro Ser
                  30
<210> 1461
<211> 54
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<220>

<221> SIGNAL

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<222> -43..-1
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<400> 1461

Met Arg Gly Trp Xaa Ala Pro Ala Trp Arg Xaa Leu Xaa Thr Arg Arg -40 -35

Leu Pro Met Gly Ser Arg His Gly Ala Ser Pro Ala Ser Ala Val Trp -20

Cys Leu Xaa Leu Lys Leu Val Pro Ala Leu Cys Ile Ser Gly Leu Thr -5

Leu Gly Ile Gln Gly Phe 10

<210> 1462

<211> 49

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -34..-1

<400> 1462

Met Tyr Phe Lys Thr Thr Xaa Xaa His Ser Ala His Met Leu Leu -25

Gln Ile Cys Phe Phe Arg Leu Thr Ile Leu Xaa Phe His Asp Asn Thr -10

Trp Gly Ser Thr Ser Phe Ser Xaa Val Ala Ala Met Leu Phe His Tyr

Arg

15

<210> 1463

<211> 26

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -24..-1

<400> 1463

Met Ser Ser Asn Ile Gln Arg Leu Gly Phe Pro Leu Leu Phe Leu Phe -20 -15

Phe Leu Phe Leu Phe Phe Phe Phe Phe

<210> 1464

<211> 69

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -67..-1

<400> 1464

Met Cys Asp Ala Phe Val Gly Thr Trp Lys Leu Val Ser Ser Glu Asn -65 -60 -55

Phe Asp Asp Tyr Met Lys Glu Val Gly Val Gly Phe Ala Thr Arg Lys -50 -45 -40

Val Ala Gly Met Ala Lys Pro Asn Met Ile Ile Ser Val Asn Gly Asp -30 -25

Val Ile Thr Ile Pro His Leu Val Leu Pro Leu Pro Met Leu Pro Thr

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WO 99/53051
                                                              PCT/IB99/00712
                                       629
                                      -10
                                                          -5
 Ser Asn Arg Lys Arg
 <210> 1465
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 <222> -21..-1
 <400> 1465
 Met Phe Leu Tyr Arg Ser Phe Gly Gly Gln Leu Leu Ser Phe Leu Leu
                         -15
                                             -10
 Gly Thr Tyr Leu Gly Arg Arg Glu Val Ala Gly Pro Gln His Gly Gln
 -5
                     1 .
                                      5
 Phe Ser Lys
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<400> 1466
 Met Xaa Gly Phe Phe Cys Leu Cys Ala Phe Asn Ser Phe Leu Leu Ser
  -15
                        -10
 Pro Glu Gly
<210> 1467
<211> 68
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<222> -66..-1
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Met Ile Phe Pro His Cys Met Tyr Cys Leu Glu Cys Ile Thr Lys Asn
                        -60
Gly Leu Leu Gly Leu Lys Val Leu Pro Leu Tyr Gly Ile Met Leu Ile
                    -45
                                        -40
Phe Phe Pro Lys Val Val Tyr Asn Asn Gln Pro Leu His Tyr Lys Ser
                -30
                                     -25
Val Met Val Phe Gln Leu Thr Ser Phe Leu Ser Ile Xaa Ile Phe Val
            -15
                                 -10
Asn Pro Thr Arg
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<213> Homo sapiens
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<210> 1469 <211> 94 <212> PRT <213> Homo sapiens <220>

<221> SIGNAL <222> -31..-1

<400> 1469

 Met
 Ala
 Ala
 Ala
 Thr
 Leu
 Thr
 Ser
 Lys
 Leu
 Tyr
 Ser
 Leu
 Leu
 Phe
 Arg

 -30
 -30
 -25
 -25
 -20
 -20

 Arg
 Thr
 Ser
 Thr
 Phe
 Ala
 Leu
 Thr
 Ile
 Xaa
 Arg
 Xaa
 Xaa
 Ser
 Cys
 Ser

 Ser
 Xaa
 Ala
 Pro
 Ser
 Ile
 Lys
 Ala
 Arg
 Thr
 Leu
 Ser
 Thr
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 Ser
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 Thr
 Ser
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<210> 1470 <211> 83 <212> PRT <213> Homo sapiens

<220>

<221> SIGNAL <222> -41..-1

<400> 1470

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 Lys
 Ala
 Ile
 Lys
 Lys
 Ser
 Leu
 Thr
 Glu
 Glu
 Glu
 Tyr
 Leu
 Tyr
 Leu

 Asp
 Phe
 Ser
 His
 Gln
 Thr
 Glu
 Gly
 Cys
 Ile
 Phe
 Pro
 Leu
 His
 Thr
 Ser

 -25
 -20
 -20
 -15
 -15
 -15
 -10
 Phe
 Lys
 Ile
 Arg
 Arg
 Arg
 Arg
 Arg
 Arg
 Arg
 Glu
 Ile
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631
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Met Phe Leu Cys Val Cys Tyr Phe Ile Arg Lys Ser Thr Ser Phe Phe
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 Ser Ile Ser Ser
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<210> 1472
<211> 71
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<222> -45..-1
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Met Gly Lys Pro Arg Gly Glu Met Leu Glu Val Val Lys Thr Val
                    -40
                                        -35
Ser Thr Phe Thr Leu Gly Gly Trp Lys Gly Thr Ala Pro Val Ser Cys
                -25
                                    -20
Ala Trp Trp Leu Leu Pro Val Trp Lys Leu Gly Gly Gln Leu Glu
           -10
                                -5
Arg Arg Lys Asn Pro Lys Glu Tyr Cys Leu Gly Ser Trp Val Trp Leu
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Ser Pro Gln Leu Ala Pro Arg
<210> 1473
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Met Leu Ile Phe Thr Phe Ile Ser Thr Leu Leu Phe Val Phe Leu Gly
   -15
                       -10
Val Val
1
<210> 1474
<211> 47
<212> PRT
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<222> -37..-1
<400> 1474
Met Glu Val Leu Ser Xaa Pro Asn Ser Phe Gln Thr Gln Ala Leu Trp
                           -30
                                               -25
Asp Ser Leu His Ser Pro Gly Val Pro Gly Ser Gly Leu Cys Ser Met
                       -15
                                           -10
Ala Ala Val Gln Ala Gly Asn Gln Ala Ile Tyr Ser Ala Ser Gly
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WO 99/53051 - 5 <210> 1475 <211> 47 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -42..-1 <400> 1475 Met Gln Ala Thr Ala Ser Gln Pro Ile His Phe Phe Xaa Ser Ser Pro -35 Gln Ala Pro Arg His His Ser Gly His Pro Val Pro Leu Leu Thr -20 -15 Gln Ala Gly Phe Pro Arg Arg Gly Glu Ala Ala Pro Pro Leu Leu -5 1 <210> 1476 <211> 34 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -30..-1 <400> 1476 Met Arg Gly Xaa Asn Xaa Val Phe Arg Val Phe Ser Glu Ser Leu Lys -25 -20 Gly Leu Cys Thr Phe Thr Leu Asn Leu Thr Ala Val Arg Thr Ile Xaa

Leu Asp

<210> 1477 <211> 40 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -32..-1

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Phe Met Glu Leu Ser Ser Leu Arg

<210> 1478 <211> 112 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -67..-1

Met Asn Leu Val Ile Cys Val Leu Leu Leu Ser Ile Trp Lys Asn Asn

633 -60 Cys Met Thr Thr Asn Gln Thr Asn Gly Ser Ser Thr Thr Gly Asp Lys -45 -40 Pro Val Glu Ser Met Gln Thr Lys Leu Asn Tyr Leu Arg Arg Asn Leu -30 -25 Leu Ile Leu Val Gly Ile Ile Ile Met Val Phe Val Phe Ile Cys Phe -15 -10 Cys Tyr Leu His Tyr Asn Cys Leu Ser Asp Asp Ala Ser Lys Ala Gly Met Val Lys Lys Gly Ile Ala Ala Lys Ser Ser Lys Thr Ser Phe 20 25 Ser Glu Ala Lys Thr Ala Ser Gln Cys Ser Ser Glu Thr Gln Thr Gly <210> 1479 <211> 35 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -28..-1 <400> 1479 Met Gln Ile Ser Ala Ala Ser Leu Asn Phe Ser Ser Lys Asn Gly Ile -25 -20 Phe Phe Ser Leu Thr Leu Ser Gly Cys Lys Phe Ser Lys Leu Leu Cys ~5 Pro Phe Gly <210> 1480 <211> 72 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -52..-1 <400> 1480 Met Ile Phe Glu Pro Val Val Leu Lys Pro Val Phe Leu Asn Ile Phe -45 Phe Phe Ser His His Val Phe Thr Val Phe Phe Ser Gly Ser His Val -30 -25 Asp Ile Leu Ser Arg Thr Val Leu Val Trp Asp Cys Leu Leu Pro Pro -15 -10 Pro Ser Phe Phe Leu Leu Leu Ser Ser Ser Xaa Ser Xaa Leu Leu Leu Xaa Xaa Ser Ser Ser Ser Arg <210> 1481 <211> 20 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -14..-1 <400> 1481

Met Leu Val Pro Leu Leu Ser His Leu Leu Phe Lys Phe Thr Trp Pro

Lys Xaa Ser Gln

5

<210> 1482

<211> 70

<212> PRT

<213> Homo sapiens

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<221> SIGNAL

<222> -49..-1

<400> 1482

Met Asp Arg Asn Pro Ser Pro Pro Pro Pro Gly Arg Asp Lys Glu Glu
-45 -40 -35

Glu Glu Glu Val Ala Gly Gly Asp Cys Ile Gly Ser Thr Val Tyr Ser
-30 -25 -20

Lys His Trp Leu Phe Gly Val Leu Ser Gly Leu Xaa Gln Xaa Val Ser
-15 -10 -5

Pro Gly Lys His Gln Asn Leu Gly Ser Xaa Xaa Glu Glu Gln Leu Thr 1 5 10 15

Glu Leu Asp Glu Arg Asn 20

<210> 1483

<211> 37

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -23..-1

<400> 1483

Met Lys Leu Ser Leu Ala Gly Tyr Glu Ile Leu Gly Cys His Phe Phe
-20 -15 -10

Ser Leu Ala Leu Leu Asn Thr Gly Pro Gln Tyr Leu Leu Ala Tyr Arg
-5 1 5

Val Ser Ala Glu Arg 10

<210> 1484

<211> 48

<212> PRT

<213> Homo sapiens

<220> ·

<221> SIGNAL

<222> -40..-1

<400> 1484

Met Ala Thr Ser Val Gly His Arg Cys Leu Gly Leu Leu His Gly Val
-40 -35 -30 -25

Ala Pro Trp Arg Ser Ser Leu His Pro Cys Glu Ile Thr Ala Leu Ser

-20 -15 -10

Gln Ser Leu Gln Pro Leu Arg Lys Leu Pro Phe Arg Ala Sar Kaa Thr
-5 1 5

<210> 1485

<211> 126

<212> PRT

<213> Homo sapiens

<213> Homo sapiens

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<220>
<221> SIGNAL
<222> -49..-1
<400> 1485
Met Ala Pro Lys Gly Lys Val Gly Thr Arg Gly Lys Lys Gln Ile Phe
               -45
                                    -40
Glu Glu Asn Arg Glu Thr Leu Lys Phe Tyr Leu Arg Ile Ile Leu Gly
           -30
                                -25
Ala Asn Ala Ile Tyr Cys Leu Val Thr Leu Val Phe Phe Tyr Ser Ser
                           -10
Ala Ser Phe Trp Ala Trp Leu Ala Leu Gly Phe Ser Leu Ala Val Tyr
Gly Ala Ser Tyr His Ser Met Ser Ser Met Ala Arg Ala Ala Phe Ser
               20
                                   25
Glu Asp Gly Ala Leu Met Asp Gly Gly Met Asp Leu Asn Met Glu Gln
                               40
Gly Met Ala Glu His Leu Lys Asp Val Ile Leu Leu Thr Ala Ile Val
                           55
Gln Val Leu Ser Cys Phe Ser Leu Tyr Val Trp Ser Phe Trp
<210> 1486
<211> 55
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -29..-1
<400> 1486
Met Ala Ala Val Thr Val Thr Lys Thr Ala Ala Ala Thr
                                -20
Ala Phe Asn Lys Ala Val Trp Phe Thr Pro Cys Ser Cys Gln Glu Val
                               ~5
Ser Ser Arg Leu Pro Ala Arg Thr Ala Ala Thr Arg Gln Asp Arg Ala
Asp Lys Lys Glu Arg Pro Cys
<210> 1487
<211> 34
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -19..-1
<400> 1487
Met Leu Gln Phe Glu Lys Pro Gly Ser Ala Ile Cys Leu Trp His Ser
               -15
                                   -10
Thr Leu Cly Gly Xaa Gly Gly Arg Glu Ile Xaa Ser Leu Arg Pro Ala
                           5
Cys Gly
    15
<210> 1488 '
<211> 24
<212> PRT
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<220>
 <221> SIGNAL
  <222> -18..-1
  <400> 1488
 Met Leu Ile Ser Tyr Leu Ala Ile Leu Leu Lys Trp Val Ser Asn Ser
           -15
                                  -10
 Lys Ser Phe Leu Val Lys Ala Ser
 <210> 1489
 <211> 76
 <212> PRT
  <213> Homo sapiens
 <220>
 <221> SIGNAL
  <222> -15..-1
 <400> 1489
 Met Lys Leu Gln Thr Leu Ala Phe Trp Ser Ala Tyr Val Pro Cys Gln
                     -10
 Thr Gln Asp Arg Asp Ala Pro Arg Leu Thr Leu Glu Gln Ile Asp Leu
 Ile Arg Arg Met Cys Ala Ser Tyr Ser Glu Leu Glu Leu Val Thr Ser
 Ala Lys Ala Leu Asn Asp Thr Gln Lys Leu Ala Cys Leu Ile Gly Val
                         40
 Glu Gly Gly His Ser Leu Asp Asn Ser Leu Ser Arg
                     55
 <210> 1490
 <211> 23
 <212> PRT
 <213> Homo sapiens
 <220>
 <221> SIGNAL
 <222> -14..-1
 <400> 1490
 Met Pro Ala Cys Leu Ser Ser Phe Val Ile Pro Ser Leu Leu Ser Pro
                 -10
 Ser Ser Pro Pro Ser Ile Gly
      5
 <210> 1491
 <211> 34
 <212> PRT
 <213> Homo sapiens
 <220>
 <221> SIGNAL
 <222> -16..-1
 :400> 1491
 Met Val Val Ser Phe Ala Gly Ser Cys Thr Ile Leu Gly Ala Ser Ser
                        -10
                                             - 5
 His Ser Phe Pro Ile Glu Val Ser Leu Phe Pro Val Asp Cys Gly Phe
                                     10
 Leu Leu
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<210> 1492
<211> 32
<212> PRT
<213> Homo sapiens
<220>
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<222> -20..-1
<400> 1492
Met Cys Cys Pro Gly Trp Asn Ala Val Ser Gln Ser Trp Leu Ala Ala
-20
                    -15
                                        -10
Pro Ser Thr Ser Trp Val Gln Glu Ile Leu Val Leu Gln Pro Pro Gly
                1
                                5
<210> 1493
<211> 69
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -54..-1
<400> 1493
Met Gly Glu Ile Lys Val Ser Pro Asp Tyr Asn Trp Phe Arg Gly Thr
               -50
                                 -45
Val Pro Leu Lys Xaa Xaa Xaa Val Asp Asp Asp Ser Lys Ile Trp
            -35
                                -30
                                                    -25
Ser Xaa Tyr Asp Ala Gly Pro Arg Ser Ile Arg Cys Pro Leu Ile Phe
     -20
                            -15
                                                -10
Leu Xaa Xaa Val Ser Gly Thr Xaa Asp Val Phe Phe Arg Gln Ile Leu
Ala Leu Thr Gly Trp
                15
<210> 1494
<211> 45
<212> PRT
<213> Homo sapiens
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<222> -16..-1
<400> 1494
Met Asp Ala Ser His Ser His Leu Ser Leu Val Gly His Ser Arg Ala
                       -10
                                            -5
Cys Gly Val Thr Ser Arg Pro His Ala Arg His Arg Gly Arg Cys Leu
              5
                                   10
Gly Pro Cys Ser Arg Ser Gly Pro Arg Leu Cys Ser Ala
            20
<210> 1495
<211> 61
-212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -34..-1
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<400> 1495

Met Gly Ser Asn Ala Val Val Trp His Thr Lys Pro Ser Leu Leu Asn -30 -25 His Pro Ala Ser Ser Leu Ile Ser His Asp Pro Trp Pro Arg Gly Ala -10 -5 Phe Ala Leu Ser Cys Pro Ser Ala Ser Phe Met Leu Phe Ser Ser Leu 10 Gln Cys Pro Phe Pro Tyr Xaa Xaa Thr Glu Cys Asn Xaa 20 <210> 1496 <211> 56 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -18..-1 <400> 1496 Met Lys Glu Asp Gly Ala Cys Leu Phe Arg Ala Val Ala Asp Gln Val -15 -10 -5 Tyr Gly Asp Gln Asp Met His Glu Val Val Arg Lys His Xaa Met Asp 10 Tyr Leu Met Lys Asn Ala Asp Tyr Phe Ser Xaa Tyr Val Thr Glu Asp 15 20 Phe Thr Tyr Ile Xaa Arg Lys <210> 1497 <211> 24 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -21..-1 <400> 1497 Met Val His Leu Ile Leu Thr Glu Val Leu Ile Met Ile Xaa Glu Ala -15 -10 Xaa Asn Val Trp Cys Gly Asp Ser <210> 1498 <211> 51 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -47..-1 <400> 1498 Met Tyr His Asn Leu Phe Ala Leu Leu Leu Ile Asp Ile His Val Val -40 -35 Leu Val Phe Tyr Cys Leu Asp Leu Leu Met Ile His Ile Phe Tyr Cys -25 -20 Lys Tyr Cys Leu Xaa Phe Gly Ile Leu Ala Ser Glu Val Tyr Ser Trp -10 -5 Asn Ile Tyr <210> 1499

<211> 44

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<212> PRT
 <213> Homo sapiens
 <220>
 <221> SIGNAL
 <222> -29..-1
 <400> 1499
 Met Glu Ser Pro Ser Arg Ala Gly Gly Val Xaa Leu Xaa Lys Ala Ala
                -25
                                     -20
 Ser Pro Leu Cys Ser Xaa Ser Ser Gly Tyr Cys Xaa Ala Phe Pro Arg
            -10
                                - 5
 Arg Ser Ala Arg Arg His Leu His Pro Gly His Gly
                        10
 <210> 1500
 <211> 61
 <212> PRT
 <213> Homo sapiens
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 <221> SIGNAL
 <222> -25..-1
<400> 1500
Met Trp Arg Tyr Val Ser Arg Leu Ser Ser Val Pro Leu Ile Ser Leu
             -20
                                   -15
Ser Val Leu Met Pro Val Gln His Ser Pro Asp Phe Cys Ser Phe Ile
                -5
Val Ser Thr Val Ile Pro Trp Phe Pro Trp Gly Ile Gly Ser Arg Thr
                           15
Leu Met Asp Ile Lys Ile Leu Gly Cys Ser Ser Pro Gly
<210> 1501
<211> 33
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -30..-1
<400> 1501
Met Asp Val Ser Cys Lys Ile Leu Tyr Asn Val Ile Glu Lys Phe Cys
                -25
                                   -20
Asn Asn Leu Leu Lys Leu Ser Ser His Ser Pro Thr Cys Ala Cys Lys
               -10
Leu
<210> 1502
<211> 29
<212> PRT
<213> Homo sapiens
<220>
<221: SIGNAL
<222> -20..-1
<400> 1502
Met Ile Phe Lys Asp Val Phe Ser His Leu Ser Gly Ser Ser Leu Gln
                   -15
                                   -10
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Leu Cys Val Ala Gln Phe Leu Xaa Leu Ser Ala Val Asp

<210> 1506 <211> 115 <212> PRT 5

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<210> 1503
 <211> 50
 <212> PRT
<213> Homo sapiens
<220>
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<222> -44..-1
<400> 1503
Met Lys Leu Thr Lys Asn Ile Leu Xaa Val Ile Ile Gly Cys Phe Lys
                -40
                                     -35
Leu Ile Ala Tyr Lys Asn Ser Val Leu Tyr Phe Tyr Ser Asn Phe Ser
            -25
                                -20
Phe Ser Phe Leu Phe Phe Phe Leu Ser Phe Phe Phe Phe Phe
Phe Phe
5
<210> 1504
<211> 92
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -87..-1
<400> 1504
Met Asn Asn Gln Lys Gln Xaa Xaa Pro Thr Leu Ser Gly Gln Arg Phe
                            -80
                                                -75
Lys Thr Arg Lys Arg Asp Glu Lys Glu Arg Phe Asp Pro Thr Gln Phe
                        -65
                                            -60
Gln Asp Cys Ile Ile Gln Gly Leu Thr Glu Thr Gly Thr Asp Leu Glu
                    -50
                                        -45
Ala Val Ala Lys Phe Leu Asp Ala Ser Gly Ala Lys Leu Asp Tyr Arg
                -35
                                    -30
                                                        -25
Arg Tyr Ala Glu Thr Leu Phe Asp Ile Leu Val Ala Gly Xaa Met Leu
            -20
                               -15
Ala Pro Gly Gly Thr Leu Ala Asp Asp Met Met Xaa
                            1
<210> 1505
<211> 35
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -17..-1
<4005 1505
Met Ala Asp Ser Leu Glu Ile Lys Leu Pro Phe Leu Pro Phe Ala Gln
                           -10
Gln Ile Asp Ile Lys Ser Cys Phe Tyr Phe Phe Yaa Asn Xaa Xaa
Phe Pro Arg
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WO 99/53051 641 <213> Homo sapiens <220> <221> SIGNAL <222> -35..-1 <400> 1506 Met Asp Arg Lys Trp Thr Trp Lys Arg Gly Gln Arg Ser His Leu Glu -30 -25 Ser Gly Gln Ala Ala Pro Ala Thr Ala Ala Ala Thr Ala Ala Ser Ala Thr Thr Gly Ala Ser Val Trp Arg Ser Thr Met Gly Xaa Leu Cys Asp Cys Thr Xaa Xaa Pro Tyr Glu Gly Pro Phe Cys Lys Lys Glu Val Ser Ala Val Phe Glu Ala Gly Thr Ser Val Thr Tyr Met Phe Gln Glu Pro 40 Tyr Pro Val Thr Lys Asn Ile Ser Leu Ser Ser Ser Ala Ile Tyr Thr 55 Asp Ser Ala Pro Ser Lys Glu Asn Ile Ala Leu Ser Phe Val Thr Thr Gln Ala Pro 80 <210> 1507 <211> 74 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -43..-1 <400> 1507 Met Ala Pro Gln Met Tyr Glu Phe His Leu Pro Leu Ser Pro Glu Glu -35 Leu Leu Lys Ser Gly Gly Val Asn Gln Tyr Val Val Gln Glu Val Leu -20 -15 Ser Ile Lys His Leu Pro Pro Gln Leu Arg Ala Phe Gln Ala Ala Phe -5 1 Arg Ala Gln Gly Pro Leu Ala Met Leu Gln His Phe Asp Thr Ile Tyr 10 Ser Ile Leu His His Phe Arg Ser Ile Asp <210> 1508 <211> 84 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -15..-1 <400> 1508 Met Ala Ala Val Gln Val Val Gly Ser Trp Pro Ser Val Gln Pro Arg -10 ·- 5 Glu Ala Pro Arg Glu Ala Ile Pro Glu Arg Gly Asn Gly Phe Arg Leu

10 Leu Ser Ala Arg Leu Cys Ala Leu Arg Pro Asp Asp Ser Ser Ser Ala 25 Arg Thr Glu Ile His Leu Xaa Phe Asp Gln Leu Ile Ser Glu Asn Tyr

642 Ser Glu Gly Ser Gly Val Ala Pro Glu Asp Val Ser Ala Leu Leu Val 60 Gln Ala Cys Gly <210> 1509 <211> 48 <212> PRT . <213> Homo sapiens <220> <221> SIGNAL <222> -30..-1 <400> 1509 Met Phe His Gly Cys His Ile Leu Ser Phe Leu Arg Ile Ser Thr Arg -25 -20 Gly Phe Leu Phe Phe Leu Gln Phe Ser Phe Pro Leu Tyr Tyr Leu Phe -10 -5 Arg Xaa Xaa Phe Pro Gln Ser Phe Met Leu Glu Ala Phe Val Arg Cys . 5 10 <210> 1510 <211> 42 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -26..-1 <400> 1510 Met Tyr Arg His Ser Lys Gln Arg Asn Asn Val Pro Cys Leu Val Leu -20 ~15 Tyr Ala Pro Trp Val Pro Pro Leu Leu Leu Ala Phe Trp Gly Trp Trp -5 Leu Leu Glu Gln Gly Leu Phe Phe Phe <210> 1511 <211> 137 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -50..-1 Met Gly Asp Pro Ser Lys Gln Asp Ile Leu Thr Ile Phe Lys Arg Leu -45 -40 Arg Ser Val Pro Thr Asn Lys Val Cys Phe Asp Cys Gly Ala Lys Asn -30 -25 Pro Ser Trp Ala Ser Ile Thr Tyr Gly Val Phe Leu Cys Ile Asp Cys -15 -10 Ser Gly Ser His Arg Ser Leu Gly Val His Leu Ser Phe Ile Arg Ser 1 5 10 Thr Glu Leu Asp Ser Asn Trp Ser Trp Phe Gln Leu Arg Cys Met Gln 25 Val Gly Gly Asn Ala Ser Ala Ser Ser Phe Phe His Gln His Gly Cys 40 Ser Thr Asn Asp Thr Asn Ala Lys Tyr Asn Ser Arg Ala Ala Gln Leu

50 55 60 Tyr Arg Glu Lys Ile Lys Ser Leu Ala Ser Gln Ala Thr Arg Lys His

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65
                             70
                                                  75
 Gly Thr Asp Leu Trp Leu Asp Ser Cys
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 <211> 26
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 <222> -22..-1
 <400> 1512
 Met Pro Leu Pro Pro Asn Gln Ser Pro Leu Leu Leu His Leu Val Phe
                             -15
                                                 -10
 His Gln Arg Thr Leu Ile Ser Leu Pro Pro
 <210> 1513
 <211> 21
 <212> PRT
 <213> Homo sapiens
 <220>
 <221> SIGNAL
 <222> -13..-1
 <400> 1513
 Met Phe Leu Thr Phe Phe Phe Cys Thr Gln Val His Gly Pro Ser Ile
      -10
 Leu Asp Ser Pro Ala
   5
 <210> 1514
 <211> 56
 <212> PRT
 <213> Homo sapiens
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 <221> SIGNAL
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 Met Val Thr Leu Trp Ile Phe Gln Phe Phe Leu Cys Leu Thr Cys Lys
                 -10
                                     -5
 Ala Tyr Asn Leu Arg Asn Cys Asn Asp Gly Lys Gly Xaa Xaa Ser Xaa
                             10
 Val Leu Gly Leu Glu Gln Xaa Leu Pro Glu Ser Ala Gly Met Val Xaa
                         25
 Phe Leu Gly Leu Lys His Arg Trp
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 <210> 1515
 <211> 37
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 <213> Homo sapiens
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Met Val Leu Trp Ala Gly Pro Xaa Val Pro Leu Leu Cys Ala Ala Xaa
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Gly Leu Gly Ala Leu His Pro Arg Cys Ser Ser Gln Gly Leu Arg Leu
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Ala Xaa Ser Glu Ala
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<210> 1516
<211> 61
<212> PRT
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<221> SIGNAL
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Met Asn Trp Arg Arg Lys Ser Val Ile Gly Leu Ser Phe Asp Phe Val
                        -35
Ala Leu Asn Leu Thr Gly Phe Val Ala Tyr Ser Val Phe Asn Ile Gly
                    -20
                                        -15
Leu Leu Trp Val Pro Xaa Xaa Xaa Gly Ala Val Ser Pro Gln Ile Pro
                -5
Gln Arg Ser Glu Pro Arg Glu Gln Gln Arg Arg Leu Leu
<210> 1517
<211> 149
<212> PRT
<213> Homo sapiens
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Met Glu Pro Leu Ala Ala Tyr Pro Leu Lys Cys Ser Gly Pro Arg Ala
Lys Val Phe Ala Val Leu Leu Ser Ile Val Leu Cys Thr Val Thr Leu
            20
Phe Leu Leu Gln Leu Lys Xaa Leu Lys Pro Lys Ile Asn Ser Phe Tyr
Ala Phe Glu Val Lys Asp Ala Lys Gly Arg Thr Val Ser Leu Glu Lys
                                            60
Tyr Lys Gly Lys Val Ser Leu Val Val Asn Val Ala Ser Asp Cys Gln
                    70
                                        75
Leu Thr Asp Arg Asn Tyr Leu Gly Leu Lys Glu Leu His Lys Glu Phe
                                    90
Gly Pro Ser His Phe Ser Val Leu Ala Phe Pro Cys Asn Gln Phe Gly
                                105
Glu Ser Glu Pro Arg Pro Ser Lys Glu Val Glu Ser Phe Ala Arg Lys
                            120
Asn Tyr Gly Val Thr Phe Pro Ile Phe His Lys Ile Lys Ile Leu Gly
                        135
Ser Glu Gly Glu Leu
145
<210> 1518
<211> 132
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Met Asn Glu Ala Met Ala Thr Asp Ser Pro Arg Arg Pro Ser Arg Cys
                                    10
Thr Gly Gly Val Val Arg Pro Gln Ala Val Thr Glu Gln Ser Tyr
                                25
Met Glu Ser Val Val Thr Phe Leu Gln Asp Val Val Pro Gln Ala Tyr
        35
                            40
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645 Ser Gly Thr Pro Leu Thr Glu Glu Lys Glu Lys Ile Val Trp Val Arg 55 Phe Glu Asn Ala Asp Leu Asn Asp Thr Ser Arg Asn Leu Glu Phe His 70 75 Glu Ile His Ser Thr Gly Ser Glu Pro Pro Leu Leu Ile Met Ile Gly 85 90 Tyr Ser Asp Gly Met Gln Val Trp Ser Ile Pro Ile Xaa Gly Glu Xaa 100 105 Lys Ser Ser Ser Leu Phe Asp Met Ala Gln Phe Glu Arg Leu Glu Ser 115 120 Cys Leu Leu His . 130 <210> 1519 <211> 46 <212> PRT <213> Homo sapiens <400> 1519 Met Pro Val Thr Arg Ala Ser Gln Pro Arg Lys Pro Ser Ser Ala Gln 10 Gln Gln Lys Ala Ala Leu Leu Xaa Asn Asn Thr Ala Leu Gln Ser Val 25 Ser Leu Arg Ser Lys Thr Thr Ile Arg Glu Arg Pro Ser Ser 40 <210> 1520 <211> 41 <212> PRT <213> Homo sapiens <400> 1520 Met Asn Gly Phe Gly Arg Leu Glu His Phe Ser Gly Ala Val Tyr Glu 10 15 Gly Gln Phe Lys Asp Asn Met Phe His Gly Leu Gly Thr Tyr Thr Phe Pro Asn Gly Ala Lys Tyr Thr Gly Ile <210> 1521 <211> 131 <212> PRT <213> Homo sapiens <400> 1521 Met Ala Lys Ile Ala Lys Thr His Glu Asp Ile Glu Ala Gln Ile Arg 10 Glu Ile Gln Gly Lys Lys Ala Ala Leu Asp Glu Ala Gln Gly Val Gly 20 25 Leu Asp Ser Thr Gly Tyr Tyr Asp Gln Glu Ile Tyr Gly Gly Ser Asp 40 Ser Arg Phe Ala Gly Tyr Val Thr Ser Ile Ala Ala Thr Glu Leu Glu 55 Asp Asp Asp Asp Tyr Ser Ser Ser Thr Ser Leu Leu Gly Gln Lys 70 75 Lys Pro Gly Tyr His Ala Pro Val Ala Leu Leu Asn Asp Ile Pro Gln 90 Ser Thr Glu Gln Tyr Asp Pro Phe Ala Glu His Arg Pro Pro Lys Ile 105 Ala Asp Arg Glu Asp Glu Tyr Lys Lys His Arg Arg Thr Met Ile Ile 115 120 Ser Gln Ser 130

<210> 1522 <211> 82

646 <212> PRT <213> Homo sapiens <400> 1522 Met Pro Ile Asn Lys Ser Glu Lys Pro Glu Ser Cys Asp Asn Val Lys 10 Val Val Val Arg Cys Arg Pro Leu Asn Glu Arg Glu Lys Ser Met Cys Tyr Lys Gln Ala Val Ser Val Asp Glu Met Arg Gly Thr Ile Thr Val His Lys Thr Asp Ser Ser Asn Glu Pro Pro Lys Thr Phe Thr Phe Asp Thr Val Phe Gly Pro Glu Ser Lys Gln Leu Asp Val Tyr Asn Leu Thr 70 Ala Arg <210> 1523 <211> 40 <212> PRT <213> Homo sapiens <400> 1523 Met Pro Asn Arg Gly Gly Asn Gly Leu Ala Pro Gly Glu Asp Arg Phe 10 Lys Pro Val Val Pro Trp Pro His Val Glu Gly Val Glu Val Asp Leu 20 25 30 Glu Ser Ile Arg Arg Ile Asn Lys <210> 1524 <211> 35 <212> PRT <213> Homo sapiens <400> 1524 Met Ser Leu Trp Leu Cys Phe Gln Cys Pro Leu Gly Val Ser Lys Ser 10 Asn Lys Lys Arg Ile Asn Leu Cys Asn Gly Phe Trp Asn Glu Lys Ile Lys Asn Arg 35 <210> 1525 <211> 47 <212> PRT <213> Homo sapiens Met Gly Thr His Val Phe Ala Ile Asn Lys Arg Thr Tyr Val Ile Ser 10 Arg Asp Arg Glu Leu Ser Thr Ala Lys Pro Xaa Cys Ser Ser Leu Leu 20 25 Thr Ala Pro Val Leu Cys Tyr Trp Arg Ala Cys Pro Leu Gln Thr 35 40 <210> 1526 <211> 56 <212> PRT <213> Homo sapiens <400> 1526 Mot Phe Cys Phe Leu Phe Ser Trp Trp Leu Arg Gly Gly Leu His Val Leu Leu Asn Thr Cys Leu Tyr Val Pro Tyr Gly Tyr Leu Ser Leu Ile Cys Leu Leu Cys Leu Trp Tyr Leu Asn Leu Tyr Lys Phe Ser Ile Phe 40

Phe Ser Phe Leu Ser Phe Phe

647 50 55 <210> 1527 <211> 55 <212> PRT <213> Homo sapiens <400> 1527 Met Thr Thr Ser Lys His Ala Ala Tyr Cys Leu Lys Gly Ser Cys Leu Xaa Gln Ala Arg Val Gln Trp Pro Leu Lys Xaa Thr Thr Ala Ser 25 Asn Phe Trp Ala Gln Val Ile Leu Ser Leu Pro Val Val Phe Val Asp Cys Leu Met Glu Xaa His Gly <210> 1528 <211> 121 <212> PRT <213> Homo sapiens <400> 1528 Met Glu Gly Gly Gly Ile Pro Leu Glu Thr Leu Lys Glu Glu Ser 10 Gln Ser Arg His Val Leu Pro Ala Ser Phe Glu Val Asn Ser Leu Gln Lys Ser Asn Trp Gly Phe Leu Leu Thr Gly Leu Val Gly Gly Thr Leu Val Ala Val Tyr Ala Val Ala Thr Pro Phe Val Thr Pro Ala Leu Arg 60 Lys Val Cys Leu Pro Phe Val Pro Ala Thr Met Lys Gln Ile Glu Asn 70 Val Val Lys Met Leu Arg Cys Arg Arg Gly Ser Leu Val Asp Ile Gly 90 Ser Gly Asp Gly Arg Ile Val Ile Ala Ala Ala Lys Lys Gly Phe Xaa 100 105 Ala Val Gly Tyr Glu Leu Asn Pro Trp <210> 1529 <211> 154 <212> PRT <213> Homo sapiens <400> 1529 Met Ala Thr Pro Leu Ala Val Asn Ser Ala Ala Ser Leu Trp Gly Pro 10 Tyr Lys Asp Ile Trp His Lys Val Gly Asn Ala Leu Trp Arg Arg Gln Pro Glu Ala Val Xaa Leu Leu Asp Lys Ile Leu Lys Lys His Lys Pro Asp Phe Ile Ser Leu Phe Lys Asn Pro Pro Lys Asn Val Gln Gln His Glu Lys Val Gln Lys Ala Ser Thr Glu Gly Val Ala Ile Gln Gly Gln

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<211> 125
<212> PRT
<213> Homo sapiens
<400> 1530
Met Asn Gly Arg Ala Asp Phe Arg Glu Pro Asn Ala Glu Val Pro Arg
Pro Ile Pro His Ile Gly Pro Asp Tyr Ile Pro Thr Glu Glu Glu Arg
Arg Val Phe Ala Glu Cys Asn Asp Glu Ser Phe Trp Phe Arg Ser Val
Pro Leu Ala Ala Thr Ser Met Leu Ile Thr Gln Gly Leu Ile Ser Lys
                        55
Gly Ile Leu Ser Ser His Pro Lys Tyr Gly Ser Ile Pro Lys Leu Ile
                    70
Leu Ala Cys Ile Met Gly Tyr Phe Ala Gly Lys Leu Ser Tyr Val Lys
Thr Cys Gln Glu Lys Phe Lys Lys Leu Glu Asn Ser Pro Leu Gly Glu
           100
                                105
Ala Leu Arg Ser Gly Gln Ala Arg Arg Ser Ser Pro Pro
        115
<210> 1531
<211> 35
<212> PRT
<213> Homo sapiens
<400> 1531
Met His Met Ser Lys Leu Ile Asn Leu Tyr Thr Ser Xaa Met Cys Asn
                                    10
Leu Leu Xaa Ile His Leu Xaa Xaa Ile Ser Cys Leu Xaa Asn Asn Lys
Xaa Thr Leu
        35
<210> 1532
<211> 111
<212> PRT
<213> Homo sapiens
<400> 1532
Met Tyr Gly Lys Gly Lys Ser Asn Ser Ser Ala Val Pro Ser Asp Ser
                                    10
Gln Ala Arg Glu Lys Leu Ala Leu Tyr Val Tyr Glu Tyr Leu Leu His
                                25
Val Gly Ala Gln Lys Ser Ala Gln Thr Phe Leu Ser Glu Ile Arg Trp
                            40
Glu Lys Asn Ile Thr Leu Gly Glu Pro Pro Gly Phe Leu His Ser Trp
                        55
Trp Cys Val Phe Trp Asp Leu Tyr Cys Ala Ala Pro Glu Arg Arg Glu
                    70
Thr Cys Glu His Ser Ser Glu Ala Lys Ala Phe His Asp Tyr Ser Ala
                                    90
Ala Ala Ala Pro Ser Pro Val Leu Gly Asn Ile Pro Pro Gly Asp
<210> 1533
<211> 107
<212> PRT
<213> Homo sapiens
<400> 1533
Met Asn Pro Glu Tyr Asp Tyr Leu Phe Lys Leu Leu Leu Ile Gly Asp
                                    10
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Ser Gly Val Gly Lys Ser Cys Leu Leu Leu Arg Phe Ala Asp Asp Thr

649 20 25 Tyr Thr Glu Ser Tyr Ile Ser Thr Ile Gly Val Asp Phe Lys Ile Arg 40 Thr Ile Glu Leu Asp Gly Lys Thr Ile Lys Leu Gln Ile Trp Asp Thr 55 Ala Gly Gln Glu Arg Phe Arg Thr Ile Thr Ser Ser Tyr Tyr Arg Gly 70 Ala His Gly Ile Ile Val Val Tyr Asp Val Thr Asp Gln Glu Ser Tyr 90 Ala Xaa Val Lys Gln Trp Leu Gln Glu Ile Asp <210> 1534 <211> 31 <212> PRT <213 > Homo sapiens <400> 1534 Met Asn Ser Lys Ala Xaa Lys Ser Ser Thr Ala Asn Gln Gly Asp Gly 10 Asp Glu Glu Xaa Val Gly Arg Xaa Glu Xaa Ser Val Gly Glu Phe <210> 1535 <211> 48 <212> PRT <213> Homo sapiens <400> 1535 Met Leu Tyr Ser Thr Leu Lys His Thr Leu Gln Tyr Val Ile Ile Asn Cys Gly His His Ala Val Gln Lys Ile Ser Lys Thr Tyr Ser Ser Cys 25 Leu Thr Glu Xaa Leu Tyr Pro Leu Pro Asn Ile Ser Pro Ile Pro Pro <210> 1536 <211> 94 <212> PRT <213> Homo sapiens <400> 1536 Met Asn Asp Glu Val Asn Pro Arg Arg Val Leu Glu Leu Met Gly Ser 10 Glu Val Thr Gln Ile Ala Cys Gly Arg Gln His Thr Leu Xaa Phe Val Pro Ser Ser Gly Leu Ile Tyr Ala Phe Gly Cys Gly Ala Arg Gly Gln Leu Gly Thr Gly His Thr Cys Asn Val Lys Cys Pro Ser Pro Val Lys Gly Tyr Trp Ala Ala His Ser Gly Gln Leu Ser Ala Arg Ala Asp Arg Phe Lys Tyr His Ile Val Lys Gln Ile Phe Ser Gly Gly Asp <210> 1537 <211> 22 <212> PRT <213> Homo sapiens <400> 1537 Met Pro Val Arg Thr Ile Thr Arg Gln Asn Gly Ser Val Pro Trp Gly

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Pro Asn His Cys Asp Lys

WO 99/53051 <211> 94 <212> PRT <400> 1538 <210> 1539 <211> 67 <212> PRT

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Glu Ile Ser Ser Ser Lys Pro Ala Ile Ser Asn Ile Leu Asn Arg Val

Asn Pro Ser Ser Tyr Ser Arg Gly Leu Lys Asn Gly Ala Leu Ser Arg 70

Gly Ile Thr Ala Ala Phe Lys Pro Thr Ser Gln His Tyr Thr

<213> Homo sapiens

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Met Val Thr Gln Ala Gln Gln Glu Ile Thr Val Gln Gln Leu Met Ala

His Leu Asp Ala Ile Arg Lys Asp Met Val Ile Leu Glu Lys Ser Glu 25

Phe Ala Asn Leu Arg Ala Glu Asn Glu Lys Met Lys Ile Glu Leu Asp

Gln Val Lys Gln Gln Leu Met His Glu Thr Ser Xaa Ile Arg Ala Asp

Asn Lys Leu

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<211> 38

<212> PRT

<213> Homo sapiens

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Ser Ile Trp Ser Phe Phe Leu Phe Tyr Gly Lys Tyr Thr Tyr Ile Arg 20

Leu Ile Leu Ser Gln Gly 35

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<211> 35

<212> PRT

<213> Homo sapiens

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Met Thr Phe Asp Leu Ser Val Phe Ser Thr Leu Ser Asp His Phe Tyr 10

Ser Ser Ser Leu Ser Asn Thr Ala Arg Asn Leu Tyr Ile Cys Leu Phe

His Ile Thr

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<212> PRT

<213> Homo sapiens

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Leu Leu Pro Pro Gly Trp Leu Asp 40

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<222> -18..-1

<400> 1555

Met Lys Leu Met Val Leu Met Leu Ala Ala Leu Leu His Cys
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Tyr Ala Asp Ser Gly Cys Lys Leu Leu Glu Asp Met Val Glu Lys Thr

Ile Asn Ser Asp Ile Ser Ile Pro Glu Tyr Lys Glu Leu Leu Gln Glu
15 20 25 30

Phe Ile Asp Ser Asp Ala Ala Ala Glu Ala Met Gly Lys Phe Lys Gln 35 40 45

Cys Phe Leu Asn Gln Ser His Arg Thr Leu Lys Asn Phe Gly Leu Met 50 55 60

Met His Thr Val Tyr Asp Ser Ile Trp Cys Asn Met Lys Ser Asn 65 70 75

<210> 1556

<211> 95

<212> PRT

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<222> -31..-1

<400> 1556

Met Val Ala Met Ala Ala Gly Pro Ser Gly Cys Leu Val Pro Ala Phe
-30 -25 -20

Gly Leu Arg Leu Leu Leu Ala Thr Val Leu Gln Ala Val Ser Ala Phe
-15 -5 1

Gly Ala Glu Phe Ser Ser Glu Ala Cys Arg Glu Leu Gly Phe Ser Ser

Asn Leu Cys Ser Ser Cys Asp Leu Leu Gly Gln Phe Asn Leu Leu 20 25 30

Gln Leu Asp Pro Asp Cys Arg Gly Cys Cys Gln Glu Glu Ala Gln Phe
35 40 45

Glu Thr Lys Lys Leu Tyr Ala Gly Ala Ile Leu Glu Val Cys Gly
50 55 60

<210> 1557

<211> 101

<212> PRT

<213> Homo sapiens

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<221> SIGNAL

<222> -32..-1

<400> 1557

Met Phe Ala Pro Ala Val Met Arg Ala Phe Arg Lys Asn Lys Thr Leu
-30 -25 -20

Gly Tyr Gly Val Pro Met Leu Leu Leu Ile Val Gly Gly Ser Phe Gly
-15 -10 -5

WO 99/53051 PCT/IB99/00712

655 Leu Arg Glu Phe Ser Gln Ile Arg Tyr Asp Ala Val Lys Ser Lys Met 10 Asp Pro Glu Leu Glu Lys Lys Leu Lys Glu Asn Lys Ile Ser Leu Glu 25 Ser Glu Tyr Glu Lys Ile Lys Asp Ser Lys Phe Asp Asp Trp Lys Asn 40 Ile Arg Gly Pro Arg Pro Trp Glu Asp Pro Asp Leu Leu Gln Gly Lys Lys Ser Arg Lys Pro <210> 1558 <211> 115 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -51..-1 <400> 1558 Met Gln Ala Gln Ala Pro Val Val Val Thr Gln Pro Gly Val Gly -45 -40 Pro Gly Pro Ala Pro Gln Asn Ser Asn Trp Gln Thr Gly Met Cys Asp -30 -25 Cys Phe Ser Asp Cys Gly Val Cys Leu Cys Gly Thr Phe Cys Phe Pro -15 -10 Cys Leu Gly Cys Gln Val Ala Ala Asp Met Asn Glu Cys Cys Leu Cys Gly Thr Ser Val Ala Met Arg Thr Leu Tyr Arg Thr Arg Tyr Gly Ile 20 Pro Gly Ser Ile Cys Asp Asp Tyr Met Ala Thr Leu Cys Cys Pro His 35 40 Cys Thr Leu Cys Gln Ile Lys Arg Asp Ile Asn Arg Arg Arg Ala Met 50 55. Arg Thr Phe <210> 1559 <211> 126 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -24..-1 <400> 1559 Met Asp Lys Ser Leu Leu Clu Leu Pro Ile Leu Leu Cys Cys Phe -20 -15 Arg Ala Leu Ser Gly Ser Leu Ser Met Arg Asn Asp Ala Val Asn Glu Ile Val Ala Val Lys Asn Asn Phe Pro Val Ile Glu Ile Val Arg Cys Arg Met Cys His Leu Gln Phe Pro Gly Glu Lys Cys Ser Arg Gly Arg Gly Ile Cys Thr Ala Thr Thr Glu Glu Ala Cys Met Val Gly Arg Met 50 Phe Lys Arg Asp Gly Asn Pro Trp Leu Thr Phe Met Gly Cys Leu Lys 65 Asn Cys Ala Asp Val Lys Gly Ile Arg Trp Ser Val Tyr Leu Val Asn 80 Phe Arg Cys Xaa Arg Ser His Asp Leu Cys Asn Glu Asp Leu

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 <222> -16..-1
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 Met Asp Leu Leu Trp Ile Leu Pro Ser Leu Trp Leu Leu Leu Gly
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 Gly Pro Ala Cys Leu Lys Thr Gln Glu His Pro Ser Cys Pro Gly Pro
                                     10
 Arg Glu Leu Glu Ala Ser Lys Val Val Leu Leu Pro Ser Cys Pro Gly
Ala Pro Gly Ser Pro Gly Glu Lys Gly Ala Pro Gly Pro Gln Gly Pro
 Pro Gly Pro Pro Gly Lys Met Gly Pro Lys Gly Glu Pro Gly Asp Pro
Val Asn Leu Leu Arg Cys Gln Glu Gly Pro Arg Asn Cys Arg Glu Leu
Leu Ser Arg Ala Pro Pro
<210> 1561
<211> 60
<212> PRT
<213> Homo sapiens
<220>
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<222> -19..-1
<400> 1561
Met Glu Ser Pro Ser Xaa Ser Ala Val Val Leu Pro Ser Thr Pro Gln
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                                    ~ 10
Ala Ser Ala Asn Pro Ser Ser Pro Tyr Thr Asn Ser Ser Arg Lys Gln
Pro Met Ser Ala Thr Leu Arg Glu Arg Leu Arg Lys Thr Arg Phe Ser
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Phe Asn Ser Ser Xaa Asn Val Val Asn Val Leu Lys
<210> 1562
<211> 97
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -16..-1
·400> 1562
Met Asp Phe Trp Leu Trp Pro Leu Tyr Phe Leu Pro Val Ser Gly Ala
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Leu Arg Ile Leu Pro Glu Val Lys Val Glu Gly Glu Leu Gly Gly Ser
Val Thr Ile Lys Cys Pro Leu Pro Glu Met His Val Arg Ile Tyr Leu
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Cys Arg Glu Met Ala Gly Ser Gly Thr Cys Gly Thr Val Val Ser Thr

Thr Asn Phe Ile Xaa Ala Glu Tyr Lys Gly Arg Val Thr Leu Arg Ala 55 60 Ile Pro Thr Gln Glu Ser Val Pro Ser Gly Gly Asn Thr Ala Asp Arg

70 75

Lys

<210> 1563

<211> 82

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<213> Homo sapiens

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<221> SIGNAL

<222> -34..-1

<400> 1563

Met Val Gly Glu Ala Gly Arg Asp Leu Arg Arg Arg Ala Val Ala -30 -25

Val Thr Ala Glu Lys Met Ala Val Leu Ala Pro Leu Ile Ala Leu Val -15 -10

Tyr Ser Xaa Pro Arg Leu Ser Arg Trp Leu Ala Gln Pro Tyr Tyr Leu 5 10

Leu Ser Xaa Leu Leu Ser Xaa Ala Phe Leu Leu Val Arg Xaa Leu Pro 20 25

Pro Leu Cys His Gly Leu Pro Thr Gln Arg Glu Xaa Gly Asn Pro Ser 35

Xaa Xaa

<210> 1564

<211> 48

<212> PRT

<213> Homo sapiens

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<221> SIGNAL

<222> -17..-1

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Met Ala Gln Leu Trp Leu Ser Cys Phe Leu Leu Pro Ala Leu Val Val -10

Ser Val Ala Ala Asn Val Ala Pro Xaa Phe Leu Ala Asn Met Thr Ser 5 10

Val Ile Leu Pro Glu Asp Cys Leu Trp Val Pro Arg Pro Ser Gly Trp 25

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<211> 105

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<213> Homo sapiens

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<222> -34..-1

<400> 1565

Het Val Gly Glu Ata Gly Arg Asp Leu Arg Arg Arg Ala Val Ala -30 -25

Val Thr Ala Glu Lys Met Ala Val Leu Ala Pro Leu Ile Ala Leu Val -15 -10

Tyr Ser Val Pro Arg Leu Ser Arg Trp Leu Ala Gln Pro Tyr Tyr Leu

Leu Ser Ala Leu Leu Ser Ala Ala Phe Leu Leu Val Arg Lys Leu Pro 15

WO 99/53051 658 Pro Leu Cys His Gly Leu Pro Thr Gln Arg Glu Xaa Gly Asn Pro Cys Asp Phe Asp Trp Arg Glu Val Glu Ile Leu Met Phe Leu Ser Ala Ile 55 Val Met Met Lys Asn Arg Arg Ser Ser <210> 1566 <211> 88 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -19..-1 <400> 1566 Met Val Ala Trp Arg Ser Ala Phe Leu Val Cys Leu Ala Phe Ser Leu -15 -10 Ala Thr Leu Val Gln Arg Gly Ser Gly Asp Phe Asp Asp Phe Asn Leu Glu Asp Ala Val Lys Glu Thr Ser Ser Val Lys Gln Pro Trp Asp His 20 Thr Thr Thr Thr Thr Asn Arg Pro Gly Thr Thr Arg Ala Pro Ala 35 Lys Pro Pro Gly Ser Gly Leu Asp Leu Ala Asp Ala Leu Asp Asp Gln Asp Asp Gly Arg Arg Asn Arg Val <210> 1567 <211> 119 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -53..-1 <400> 1567 Met Ala Asp Pro Asp Pro Arg Tyr Pro Arg Ser Ser Ile Glu Asp Asp -50 -45 Phe Asn Tyr Gly Ser Ser Val Ala Ser Ala Thr Val His Ile Arg Met -30 -25 Ala Phe Leu Arg Lys Val Tyr Ser Ile Leu Ser Leu Gln Val Leu Leu -10 Thr Thr Val Thr Ser Thr Val Phe Leu Tyr Phe Glu Ser Val Arg Thr 5 Phe Val His Glu Ser Pro Ala Leu Ile Leu Leu Phe Ala Leu Gly Ser 20 Leu Gly Leu Ile Phe Ala Leu Xaa Leu Asn Arg His Lys Tyr Pro Leu 35 Asn Leu Tyr Leu Leu Phe Gly Phe Thr Leu Leu Glu Ala Leu Thr Val Ala Val Val Thr Val Leu

<210> 1568 <211> 104

<212> PRT

<213> Homo sapiens

<221> SIGNAL <222> -55..-1

<400> 1568

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 Asn
 Val
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 Arg
 Ser
 Arg
 Pro
 Gln
 Lys
 His

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10 15 20 25

Gly Leu Tyr Ile Cys Leu Asn Glu Gln Thr Gly Ser Ile Leu Leu Glu
30 35

Lys Lys Met Leu Leu Ser Val Ser

<210> 1569

<211> 126

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -62..-1

<400> 1569

Met Arg Asn Lys Lys Ile Leu Lys Glu Asp Glu Leu Leu Ser Glu Thr
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-55
-50

Gln Gln Ala Ala Phe His Gln Ile Ala Met Glu Pro Phe Glu Ile Asn
-45
-40
-35

Val Pro Lys Pro Lys Arg Arg Asn Gly Val Asn Phe Ser Leu Ala Val
-30
-25
-20
-15

Val Val Ile Tyr Leu Ile Leu Leu Thr Ala Gly Ala Gly Leu Leu Val

Val Gln Val Leu Asn Leu Gln Ala Arg Leu Arg Val Leu Glu Met Tyr

5 10 15

Phe Leu Asn Asp Thr Leu Ala Ala Glu Asp Ser Pro Ser Phe Ser Leu 20 25 30
Leu Gln Ser Ala His Pro Gly Gly His Leu Ala Glu Asp Ser Pro Ser Phe Ser Leu

Leu Gln Ser Ala His Pro Gly Glu His Leu Ala Gln Gly Ala Ser Arg

40

45

50

Leu Gln Ser Cys Arg Pro Asn Ser Pro Gly Ser Ala Ser Xaa 55 60

<210> 1570

<211> 134

<212> PRT

<213> Homo sapiens

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-55 -50 -45

Arg Leu Gln Ala Leu Arg Lys Glu Lys Ser Arg Asp Ala Ala Arg Ser
-40 -35 -30

Arg Arg Gly Lys Glu Asn Phe Glu Phe Tyr Glu Leu Ala Lys Leu Leu
-20
-15
-10

Pro Leu Pro Ala Ala Ile Thr Ser Gln Leu Asp Lys Ala Ser Ile Ile

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-5
 Arg Leu Thr Ile Ser Tyr Leu Lys Met Arg Asp Phe Ala Asn Gln Gly
                         15
                                              20
 Asp Pro Pro Trp Asn Leu Arg Met Glu Gly Pro Pro Pro Asn Thr Ser
 Val Lys Val Ile Gly Ala Gln Arg Arg Arg Ser Pro Ser Ala Leu Ala
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                                      50
 Ile Glu Val Phe Glu Ala His Leu Gly Ser His Ile Leu Gln Ser Trp
                                 65
 Met Ala Leu Tyr Leu His
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 <211> 28
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                    -15
Val Thr Thr Ala His Ala Trp Leu Thr Val Leu Val
<210> 1572
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<221> SIGNAL
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                    -15
Val Thr Thr Ala His Ala Trp Leu Thr Val Leu Val
<210> 1573
<211> 47
<212> PRT
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Met Val Gly Arg Val Arg Val Cys Arg Lys Tyr Pro Pro Thr Thr Leu
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                                        -35
Trp Glu Gly Ala Arg Gly His Arg Gln Ile Ser Val Ser Pro Trp Asn
                -25
                                    -20
The Cys Cys Ala Ala Ala Ala Ala Ala Ala Gly Ser Arg The
<210> 1574
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<212> PRT
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                              -45
 Ile Glu Arg Leu Gly Thr Pro Gln Gln Ile Ala Ile Ala Arg Glu Gly
                         -30
                                              -25
 Asp Leu Leu Thr Lys Glu Arg Leu Cys Cys Gly Leu Ser Met Phe Glu
                     -15
                                          -10
 Val Ile Leu Thr Arg Ile Arg Ser Tyr Leu Gln Asp Pro Ile Trp Arg
 Gly Pro Pro Pro Thr Asn Gly Val Met His Val Asp Glu Cys Val Glu
                             20
 Phe His Arg Leu Trp Ser Ala Met Gln Phe Val Tyr Cys Ile Pro Val
 Gly Thr Asn Glu Phe Thr Ala Glu Gln Cys Phe Gly Asp Gly Leu Asn
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 Trp Ala Gly Ser Pro Xaa Leu Ser Cys Xaa Ala Ser Ser Val Ala Leu
 Thr Cys Ser Thr Ser Val Thr Thr Cys
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Met Ala Leu Val Pro Cys Gln Val Leu Arg Met Ala Ile Leu Leu Ser
                        -65
                                             -60
Tyr Cys Ser Ile Leu Cys Asn Tyr Lys Ala Ile Glu Met Pro Ser His
                    -50
Gln Thr Tyr Gly Gly Ser Trp Lys Phe Leu Thr Phe Ile Asp Leu Val
                -35
                                     -30
Ile Gln Ala Val Phe Phe Gly Ile Cys Val Leu Xaa Asp Leu Ser Ser
                                 -15
Leu Leu Thr Arg Gly Ser Gly Asn Gln Glu Gln Glu Arg Gln Leu Lys
Lys Leu Ile Ser Leu Arg Asp Trp Met Leu Ala Val Leu Ala Phe Leu
                    15
Leu Gly Phe Leu Leu
<210> 1576
<211> 79
<212> PRT
<213> Homo sapiens
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<221> SIGNAL
<222> -69..-1
<400> 1576
Met Ala Thr His His Leu Gly Leu Pro Ala Ser Gln Pro Leu Pro Gly
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-60

Ile Leu Ser Arg Ala Pro Ser Leu Pro Pro Arg Ser Pro Ala Thr Arg

-65

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-45
 Ser Arg Val Ser Ser Pro Trp Gly Glu Ser Ser Ser Leu Leu Phe
                            -30
                                                 -25
 Pro Asp Cys His Ile Ser Phe Pro Ala Leu Thr Gly Ser Gln Leu Leu
                         -15
 Gly Asp Thr Ile Pro Arg Pro His Leu Pro Pro Thr Ala Ala Cys
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 <211> 108
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 <213> Homo sapiens
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Met Thr Pro Ser Arg Leu Pro Trp Leu Leu Ser Trp Val Ser Ala Thr
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Ala Trp Arg Ala Ala Arg Ser Pro Leu Leu Cys His Ser Leu Arg Lys
Thr Ser Ser Ser Gln Gly Gly Lys Ser Glu Leu Val Lys Gln Ser Leu
Lys Lys Pro Lys Leu Pro Glu Gly Arg Phe Asp Ala Pro Glu Asp Ser
                        20
His Leu Glu Lys Glu Pro Leu Glu Lys Phe Pro Asp Asp Val Xaa Pro
                                         40
Val Thr Lys Glu Lys Gly Gly Pro Arg Gly Pro Glu Pro Thr Arg Tyr
Gly Asp Trp Glu Arg Lys Gly Arg Cys Ile Asp Phe
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<211> 81
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<213> Homo sapiens
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<222> -51..-1
<400> 1578
Met Glu Lys Leu Arg Arg Val Leu Ser Gly Gln Asp Asp Glu Glu Gln
                        -45
                                            -40
Gly Leu Thr Ala Gln Val Leu Asp Ala Ser Ser Leu Ser Phe Asn Thr
                    -30
                                        -25
Arg Leu Lys Trp Phe Ala Ile Cys Phe Val Cys Gly Val Phe Phe Ser
                -15
                                    -10
Ile Leu Gly Thr Gly Leu Leu Trp Leu Pro Gly Gly Ile Lys Leu Phe
Ala Val Phe Tyr Thr Leu Gly Asn Leu Ala Ala Leu Xaa Val His Ala
Xaa
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<222> -93..-1

<400> 1579

Met Cys Glu Asn Gln Glu Glu Pro Ala Gly Ser Val Cys Cys His Arg -85 Val Ser Ala Cys Arg Gly Gly Thr Pro Gly Gly Gly Arg Gly Gln Ser -70 His Cys Arg Gly Pro Asp Trp Glu Asn Asn Asp Met Ala Gly Ala Ser -55 Leu Gly Ala Arg Phe Tyr Arg Gln Ile Lys Arg His Pro Gly Ile Ile -40 Pro Met Ile Gly Leu Ile Cys Leu Gly Met Gly Ser Ala Ala Leu Tyr -25 Leu Leu Arg Leu Ala Leu Arg Ser Pro Asp Val Trp Leu Gly Gln Lys -5 Glu Gln Pro Gly Ala Leu Glu Pro Pro Glu Pro Gln

<210> 1580

<211> 134

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -16..-1

<400> 1580

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105

Pro Glu Arg Leu Asp Arg 115

<210> 1581

<211> 64

<212> PRT

<213> Homo sapiens

<400> 1581

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Cys Arg Lys His Asp Asp Cys Pro Asn Lys Tyr Gly Glu Lys Lys Thr 25

Lys Glu Lys Trp Asn Leu Tnr Val His Tyr Tyr Cys Leu Leu Met Ser 40

Ser Gly Ile Trp Gln Arg Gly Lys Glu Glu Glu Gly Val Met Val Phe

<210> 1582

<211> 79

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40

35

WO 99/53051 PCT/IB99/00712

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665
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 Trp Cys Val Phe Trp Asp Leu Tyr Cys Ala Ala Pro Glu Arg Arg Glu
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Thr Glu
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Ala Pro Phe Ala Gln Arg Ile Asp Pro Thr Arg Glu Lys Leu Thr Pro
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Ser Tyr His Gly Gly Glu Pro Gly Thr Ser
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Lys Asn Cys Asp Pro Lys Cys Glu Gln Lys Cys Glu Ser Lys Cys Gln
            20
                               25
Pro Ser Cys Leu Lys Lys Leu Leu Gln Arg Cys Phe Glu Lys Cys Pro
Trp Glu Lys Cys Pro Ala Pro Pro Lys Cys Leu Pro Cys Pro Ser Gln
                        55
Ser Pro Ser Ser Cys Pro Pro Gln Pro Cys Thr Lys Pro Cys Pro Pro
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90

85

10

25

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<211> 157

<212> PRT

<213> Homo sapiens

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<210> 1593 <211> 119

<212> PRT

<213> Homo sapiens

<400> 1593

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Ser Gly Leu Thr Lys Xaa Ile Leu Gly Ser Ile Gly Ser Ala Ile Ala 100 105 110

Ala Val Ile Ala Arg Phe Tyr 115

667 <211> 81 <212> PRT <213> Homo sapiens <400> 1594 Met Tyr Ile Gln Cys Cys Glu Trp Leu Gln Ser Trp Arg Ser Lys Asp Glu Phe Cys Leu Glu Glu Ser Gly Lys Ala Ser Trp Arg Arg Glu Gln 20 Trp His Gly Pro Xaa Xaa Val Arg Ser Phe Gln Phe Ile Pro Phe Lys His Cys Ser His Val Ala Phe Lys His Ser Ile Val Leu Ala Val Thr Gln Ala His Ser Ala Lys Gly Ser Thr Ser Phe Ser Ala Met Arg Thr Tyr <210> 1595 <211> 65 <212> PRT <213> Homo sapiens <400> 1595 Met Val Gly Val Ser Val Cys His His Ile Arg Val Gly Ile Lys Arg Arg Lys Ala Ala Leu Leu Glu Leu Cys Gly Leu Leu Gln Val Arg Val 20 Ala Gly Asn Arg Thr Thr Leu Leu Leu Glu Glu Lys Arg Asn Ser Phe Ser Ala Xaa Thr Arg Lys Ala Val Phe Phe Ser Gly Asp Leu His Phe 55 Ser 65 <210> 1596 <211> 111 <212> PRT <213> Homo sapiens <400> 1596 Met Pro Ser Arg Thr Ala Arg Tyr Ala Arg Tyr Ser Pro Arg Gln Arg 10 Arg Arg Arg Met Leu Ala Asp Arg Ser Val Arg Phe Pro Asn Asp Val Leu Phe Leu Asp His Ile Arg Gln Gly Asp Leu Glu Gln Val Gly Arg Phe Ile Arg Thr Arg Lys Val Ser Leu Ala Thr Ile His Pro Ser Gly Leu Ala Ala Leu His Glu Ala Val Leu Ser Gly Asn Leu Glu Cys Val 75 Lys Leu Leu Val Lys Tyr Gly Ala Asp Ile His Gln Arg Asp Glu Ala 90 Gly Trp Thr Pro Leu His Ile Ala Cys Ser Asp Gly Tyr Leu Thr 105 <210> 1597 <211> 33 <212> PRT <213> Homo sapiens <400× 1597 Met Ala Trp Gly Gly Trp Gly Ala His Ser Ala Cys Ser Glu Glu Arg 10 Ala Thr Arg Pro Val Glu Gly Ala Tyr Ser Gly Arg Trp Gly Gln Ala

Gln

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 Phe Val Phe Pro Leu Ser His Leu His Leu Glu Ser Gln Arg Pro Pro
 Ile Gly Ser Ile Ser Ser Met Glu Val Asn Val Asp Thr Leu Glu Gln
 Val Glu Leu Ile Asp Leu Gly Asp Pro Asp Ala Ala Asp Val Phe Leu
 65
 Pro Cys Glu Asp Pro Pro Pro Thr Pro Gln Ser Ser Gly Val Asp Asn
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His Leu Glu Glu Leu Ser Leu Pro Xaa Ala Tyr Ile Arg Gln Asp His
                                 105
 Ile
 <210> 1599
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 <212> PRT
 <213> Homo sapiens
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Val Val Pro Val Gly Thr Glu Glu Ala Pro Lys Asp Thr Lys Tyr Ile
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Lys Leu Pro Val Gln Thr Leu Leu Gln Gly
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                         55
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                                                                       120
tactageeta ttaettttta gteeattggg aateactaaa aaaagtagag getttagett
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catteetegg etgettaaat catattgtaa tgttttaaat tgttatgteg teetgtataa
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ccttagg
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tgtcaccttt ccataaatac tccattccct tttgtgattt tgttctttgc acatgttgtt
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ctatetetge etggaatgtg lieteeacet tittgattgte tgeca
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120

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669
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 ccatgtgaac ataaagat
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                                                                        120
gctgagtgct tgaggacgtg tttcaacaga tggttggggt tagtgtgtgt catcacattc
                                                                        180
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tccctatggg atttgtcatt ggatgttatt tagacagaaa gagtgatgaa cggctaactg
                                                                       180
ccttccggaa caagagtatg ttatttaaaa gggaattgca acccagtgaa gaagttacct
                                                                       240
ggaagtaaag actggctaga ttatcgaatg ttcacatttt aaagttctga ga
                                                                       292
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<212> DNA
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                                                                       120
gctcccttct tgcagaaggg agggggaaac atacatttat tcatgccagt ctgttgcatg
                                                                       180
caggettttt ggetteetae ettgeaacaa aataattgea ceaacteett agtgeegatt
                                                                       240
ccgcccacag agagtcctgg arccacagtc ttttttgctt tgcattgtag gagagggact
                                                                       300
aagtgctaga gactatgtcg ctttcctgag ctaccgagag cgctcgtgaa ctggaat
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aactggggga gctgccaagc tggatcttga tgcggractt cagtcctagt ggcattttcg
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gagegtttea astaggstac aceggtacta caacaagtae atcaatgtga agaaggggag
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attccagccc ttagtcaggt tctttccagt gtcctcaaac acagtaagga gagtgctcta
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tgccgcagga cctgcagcat cccagaggtg cagattttaa tttcagtgac tgaattaaaa
                                                                       180
ggtgtcaaga agctcgaatg gtatgtaggt ctcccatggt atttcaattt aaaaagaagt
                                                                       240
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                                                                       120
aaatgettat ageteaaaca geteettgga anttaageta cacagaetgt attttattag
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                                                                       240
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<212> DNA
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                                                                       120
cccttctgga gtgcatatgt gcctttacag ttctgtttgc aaacgctgtc tagcatacta
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agaggatgtt agcaaa
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                                                                      120
toggocagag gottatttat tgaogggaot gtttootttg goocacgoga ogtagottot
                                                                      180
gttgtccttg actgggcgcc gcctcccgcc ccgccgcctc ggaagccc
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                                                                       120
aagecagagg tgttecacce caateettea ceetcaccee acateatggt ggeecetggg
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gctactttct gaatcctaaa gcgctcttcc agctttcaca tttgattccg tggcagaagg
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                                                                       120
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tegeacgeec cae
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i

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CZIS> HOIIIO	saprens					
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and,	3, C OL C					
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-210- 1622						

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caactgctgg	gaaaaaaata	aaacaccaac	cccaaccgtc	agcaacaagg	taasmgaggg	360
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